

FIG. 1

Sequence of human APRIL (SEQ ID NOS: 1 and 2)

Human G70 cDNA (SEQ ID NO 1)

Length: 1465 bp

1 GCCAACCTTC CCTCCCCCAA CCCTGGGGGCC GCCCCAGGGT TCCTGCGCAC
51 TGCCTGTTCC TCCTGGGTGT CACTGGCAGC CCTGTCCTTC CTAGAGGGAC
101 TGGAACCTAA TTCTCCTGAG GCTGAGGGAG GGTGGAGGGT CTCAAGGCAA
151 CGCTGGCCCC ACGACGGAGT GCCAGGAGCA CTAACAGTAC CCTTAGCTTG
201 CTTTCCTCCT CCCTCCTTTT TATTTTCAAG TTCCTTTTTC TTTCTCCTTG
251 CGTAACAACC TTCTTCCCTT CTGCACCACT GCCCGTACCC TTACCCGCCC
301 CGCCACCTCC TTGCTACCCC ACTCTTGAAA CCACAGCTGT TGGCAGGGTC
351 CCCAGCTCAT GCCAGCCTCA TCTCCTTTCT TGCTAGCCCC CAAAGGGCCT
401 CCAGGCAACA TGGGGGGGCC AGTCAGAGAG CCGGCACTCT CAGTTGCCCT
451 CTGGTTGAGT TGGGGGGCAG CTCTGGGGGC CGTGGCTTGT GCCATGGCTC
501 TGCTGACCCA ACAACAGAG CTGCAGAGCC TCAGGAGAGA GGTGAGCCGG
551 CTGCAGGGGA CAGGAGGCC CTCCCAGAAAT GGGGAAGGGT ATCCCTGGCA
601 GAGTCTCCCG GAGCAGAGTT CCGATGCCCT GGAAGCCTGG GAGAGTGGGG
651 AGAGATCCCG GAAAAGGAGA GCAGTGCTCA CCCAAAAACA GAAGAAGCAG
701 CACTCTGTCC TGCACCTGGT TCCCATTAAAC GCCACCTCCA AGGATGACTC
751 CGATGTGACA GAGGTGATGT GGCAACCAGC TCTTAGGCGT GGGAGAGGCC
801 TACAGGCCCA AGGATATGGT GTCCGAATCC AGGATGCTGG AGTTTATCTG
851 CTGTATAGCC AGGTCCTGTT TCAAGACGTG ACTTTCACCA TGGGTCAGGT
901 GGTGTCTCGA GAAGGCCAAG GAAGGCAGGA GACTCTATTC CGATGTATAA
951 GAAGTATGCC CTCCCACCCG GACCGGGCCT ACAACAGCTG CTATAGCGCA
1001 GGTGTCTTCC ATTTACACCA AGGGGATATT CTGAGTGTCA TAATTCCCCG
1051 GGCAAGGGCG AAACCTTAACC TCTCTCCACA TGGAACCTTC CTGGGGTTTG
1101 TGAAACTGTG ATTGTGTTAT AAAAAGTGGC TCCCAGCTTG GAAGACCAGG
1151 GTGGGTACAT ACTGGAGACA GCCAAGAGCT GAGTATATAA AGGAGAGGGA
1201 ATGTGCAGGA ACAGAGGCGT CTTCTGGGT TTGGCTCCCC GTTCCTCACT
1251 TTTCCCTTTT CATTCCCACC CCCTAGACTT TGATTTTACG GATATCTTGC
1301 TTCTGTTCCC CATGGAGCTC CGAATTCTTG CGTGTGTGTA GATGAGGGGC
1351 GGGGACGGG CGCCAGGCAT TGTTTCAGACC TGGTCGGGGC CCACTGGAAG
1401 CATCCAGAAC AGCACCACCA TCTAACGGCC GCTCGAGGGA AGCACCCGGC
1451 GGTTTGGGCG AAGTC

The proposed transmembrane domains are boxed

human G70 protein sequence (SEQ ID NO 2)

1 MPASSPFLLA PKGPPGNMGG PVREPALSVA LWLSWGAALG AVACAMALLT
51 QQTELQSLRR EVSRLQGTGG PSQNGEGYPW QSLPEQSSDA LEAWESGERS
101 RKRRAVLTQK QKKQHSLVHL VPINATSKDD SDVTEVMWQP ALRRGRGLQA
151 QGYGVRIQDA GYLLYSQVL FQDVFTMGQ VVSREGQGRQ ETLFR CIRSM
201 PSHPDRAYNS CYSAGVFHLH QGDILSVIIP RARAKLNLSP HGTF LGFV

FIG. 2A

Sequence of mouse G70 (SEQ ID NOS: 3 and 4)

Mouse G70 (SEQ ID NO 3)

1	CATGCCGAGT	GCTTTGTGTG	TGTTACCTGC	TCTAAGAAGC	TGGCTGGGCA
51	GCGTTTCACC	GCTGTGGAGG	ACCAGTATTA	CTGCGTGGAT	TGCTACAAGA
101	ACTTTGTGGC	CAAGAAGTGT	GCTGGATGCA	AGAACCCCAT	CACTGGGTTT
151	GGTAAAGGCT	CCAGTGTGGT	GGCCTATGAA	GGACAATCCT	GGCACGACTA
201	CTGCTTCCAC	TGCAAAAAAT	GCTCCGTGAA	TCTGGCCAAC	AAGCGCTTTG
251	TATTTTCATAA	TGAGCAGGTG	TATTGCCCTG	ACTGTGCCAA	AAAGCTGTAA
301	CTTGACGGCT	GCCCTGTCCT	TCCTAGATAA	TGGCACCAAA	TTCTCCTGAG
351	GCTAGGGGGG	AAGGAGTGTC	AGAGTGTAC	TAGCTCGACC	CTGGGGACAA
401	GGGGGACTAA	TAGTACCCTA	GCTTGATTTT	TTCTATTCT	CAAGTTCCTT
451	TTTATTTCTC	CCTTGCGTAA	CCCGCTCTTC	CCTTCTGTGC	CTTTGCCTGT
501	ATTCCCACCC	TCCCTGCTAC	CTCTTGGCCA	CCTCACTTCT	GAGACCACAG
551	CTGTTGGCAG	GGTCCCTAGC	TCATGCCAGC	CTCATCTCCA	GGCCACATGG
601	GGGGCTCAGT	CAGAGAGCCA	GCCCTTTCGG	TTGCTCTTTG	GTTGAGTTGG
651	GGGGCAGTTC	TGGGGGCTGT	GACTTGTGCT	GTCGCACTAC	TGATCCAACA
701	GACAGAGCTG	CAAAGCCTAA	GGCGGGAGGT	GAGCCGGCTG	CAGCGGAGTG
751	GAGGGCCTTC	CCAGAAGCAG	GGAGAGCGCC	CATGGCAGAG	CCTCTGGGAG
801	CAGAGTCCTG	ATGTCCTGGA	AGCCTGGAAG	GATGGGGCGA	AATCTCGGAG
851	AAGGAGAGCA	GTA ¹ CTCACCC	AGAAGCACAA	GAAGAAGCAC	TCAGTCCTGC
901	ATCTTGTTCC	AGTTAACATT	ACCTCCAAGG	ACTCTGACGT	GACAGAGGTG
951	ATGTGGCAAC	CAGTACTTAG	GCGTGGGAGA	GGCCTGGAGG	CCCAGGGAGA
1001	CATTGTACGA	GTCTGGGACA	CTGGAATTTA	TCTGCTCTAT	AGTCAGGTCC
1051	TGTTTCATGA	TGTGACTTTC	ACAATGGGTC	AGGTGGTATC	TCGGGAAGGA
1101	CAAGGGAGAA	GAGAAACTCT	ATTCCGATGT	ATCAGAAGTA	TGCCTTCTGA
1151	TCCTGACCGT	GCCTACAATA	GCTGCTACAG	TGCAGGTGTC	TTTCATTTAC
1201	ATCAAGGGGA	TATTATCACT	GTCAAAATTC	CACGGGCAAA	CGCAAAACTT
1251	AGCCTTTCTC	CGCATGGAAC	ATTCCTGGGG	TTTGTGAAAC	TATGATTGTT
1301	ATAAAGGGGG	TGGGGATTTT	CCATTCCAAA	AACTGGCTAG	ACAAAGGACA
1351	AGGAACGGTC	AAGAACAGCT	CTCCATGGCT	TTGCCTTGAC	TGTTGTTCTT
1401	CCCTTTGCCT	TTCCCGCTCC	CACTATCTGG	GCTTTGACTC	CATGGATATT
1451	AAAAAAGTAG	AATATTTTGT	GTTTATCTCC	CAAAAA	

FIG. 2B

Mouse G70 Length: 241 (SEQ ID NO 4)

1 MPASSPGHMG GSVREPALSV ALWLSWGAVL GAVTCAVALL IQQTELQSLR
51 REVSRLQRSG GPSQKQGERP WQSLWEQSPD VLEAWKDGAK SRRRRAVLTQ
101 KHKKKHSLVH LVPVNITSKD SDVTEVMWQP VLRRGRGLEA QGDIVRVWDT
151 GIYLLYSQVL FHDVTFTMGQ VVSREGQGRR ETLFRCIRSM PSDPDRAVNS
201 CYSAGVFHLH QGDIITVKIP RANAKLSLSP HGTFGLGFVKL *

G-70 FLAG des92 (smuG70) Strain #4081 (SEQ ID NO 19):

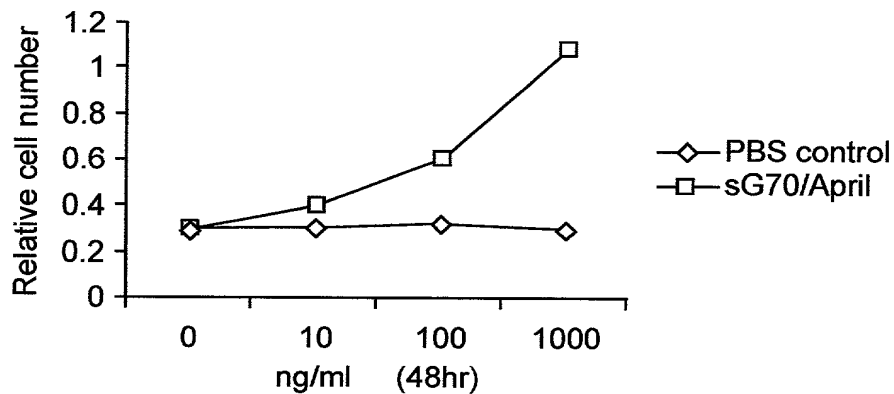
MDYKDDDDKKHKKHSLVHLVLPVNITSKDS
SDVTEVMWQPVLRGRGLEAQGDIVRVWDTGIY
LLYSQVLFHDVTFTMGQVVSREGQGRRETLFRCIRSMPSDPDRAYNSCYSAGVFHLHQGDII
TVKIPRANAKLSLSPHGTFGLGFVKL*

Alignm. of human and mouse G70

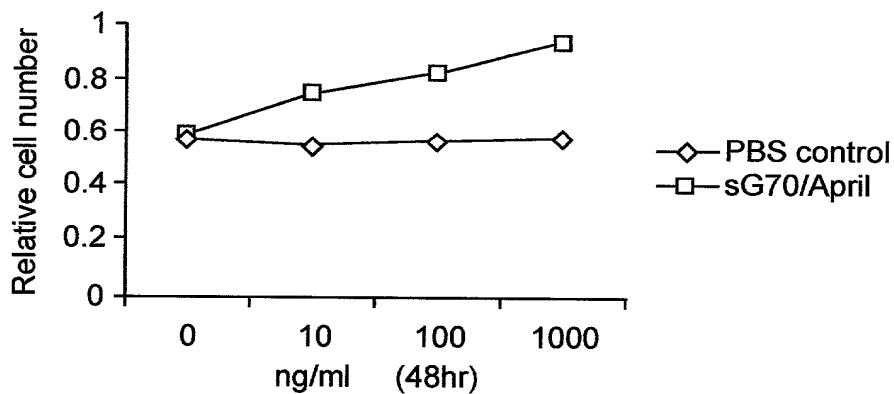
mouse:	1	MPASS-----PGHMGGS	VREPALSVALWLSWGAVLGA	VTCAVALL	IQQTE	QSLRR	51
		MPASS	PG+MGG	VREPALSVALWLSWGA	LGAV	CA+ALL	QQTE
human:	1	MPASS	FLLAPKGP	PGNMGGP	VREPALSVALWLSWGAAL	GAVACAMALL	TQQTE
							60
mouse:	52	EVSRLQ	RS	GGPSQ	KQGERPWQ	SLWEQSPDVLEAWKDGAKSR	RRRAVLTQKHKKH
		EVSRLQ	+GGPSQ	PWQSL	EQS	D LEAW+ G +SR+RR	AVLTQK
human:	61	EVSRLQ	GTG	GPSQNGEGYPWQ	SLPEQSSDALEAWESGERSR	KRRRAVLTQKQ	KKQHSVLHL
							120
mouse:	112	VPVNIT	SKD-SDVTEVMWQ	PVLR	RGRGLEAQ	GDIVRVWDTGIYLLYSQ	VLFDVTFTMGQ
		VP+N	TSKD	SDVTEVMWQ	P LRRGRGL+AQG	VR+ D G+YLLYSQ	VLFDVTFTMGQ
human:	121	VPINAT	SKDDSDVTEVMWQ	PALRRGRGLQ	AQGYGVRIQDAGVYLLYSQ	VLFDVTFTMGQ	180
mouse:	171	VVSREGQ	RRRETL	FR	CIRSMPSDPDRAYNSCYSAGV	FHLHQGDIITVKIPRANAKLS	LSP
		VVSREGQ	GR+ETL	FR	CIRSMPS PDRAYNSCYSAGV	FHLHQGDI++V IPRA AKL+LSP	
human:	181	VVSREGQ	GRQETL	FR	CIRSMPSHPDRAYNSCYSAGV	FHLHQGDIILSVIIPRARA	KNLSP
							240
mouse:	231	HGTF	LGEV	KL	240		
		HGTF	LGEV	KL			
human:	241	HGTF	LGEV	KL	250		

FIG. 4A

Effect of sG70/April on Raji cell proliferation



Effect of sG70/April on Jurkat cell proliferation



Effect of sG70/April on K562 cell proliferation

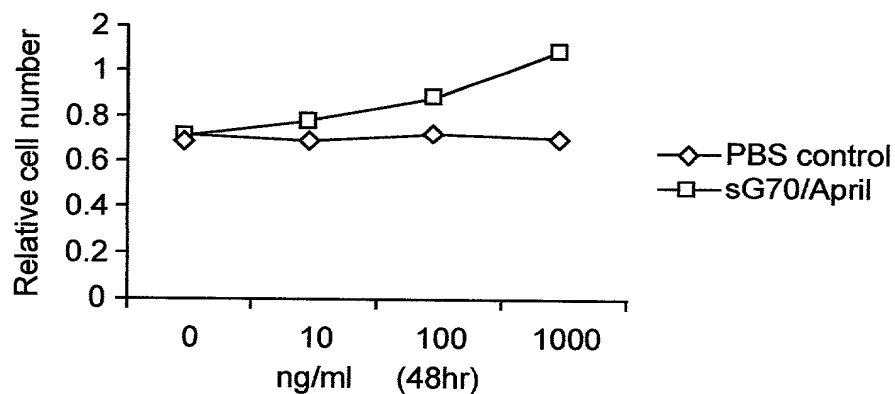


FIG. 4B

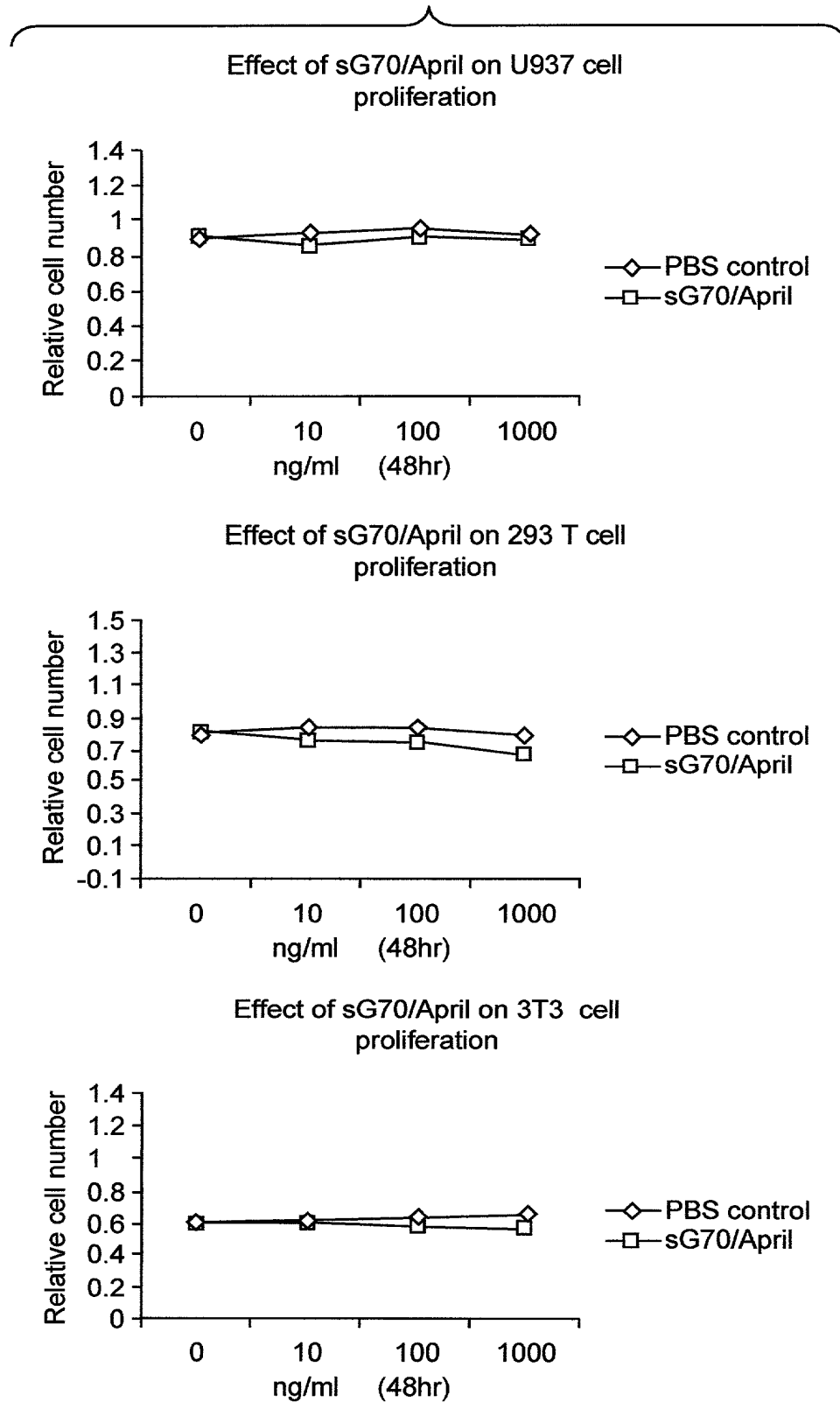


FIG. 5A

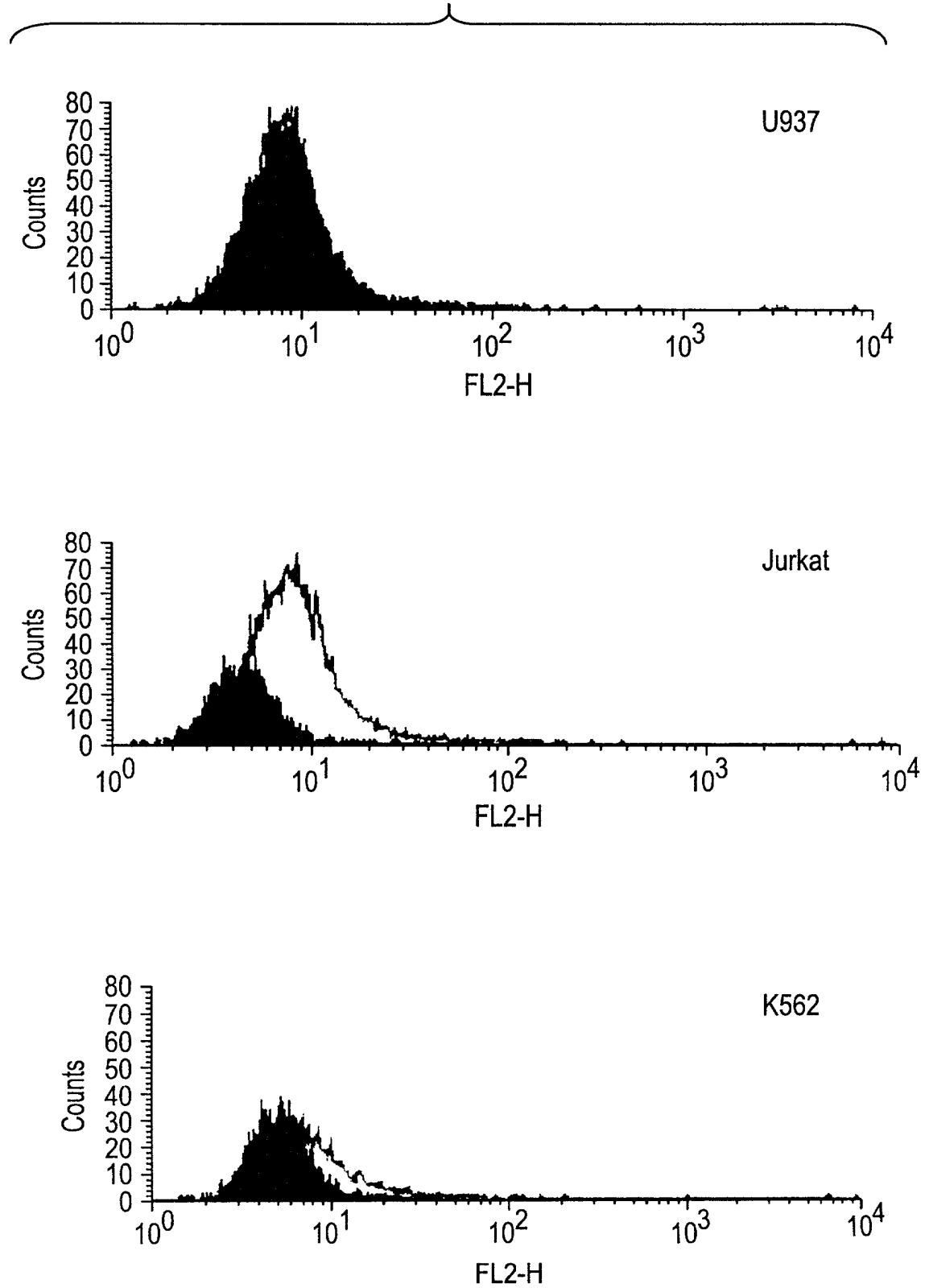


FIG. 5B-1

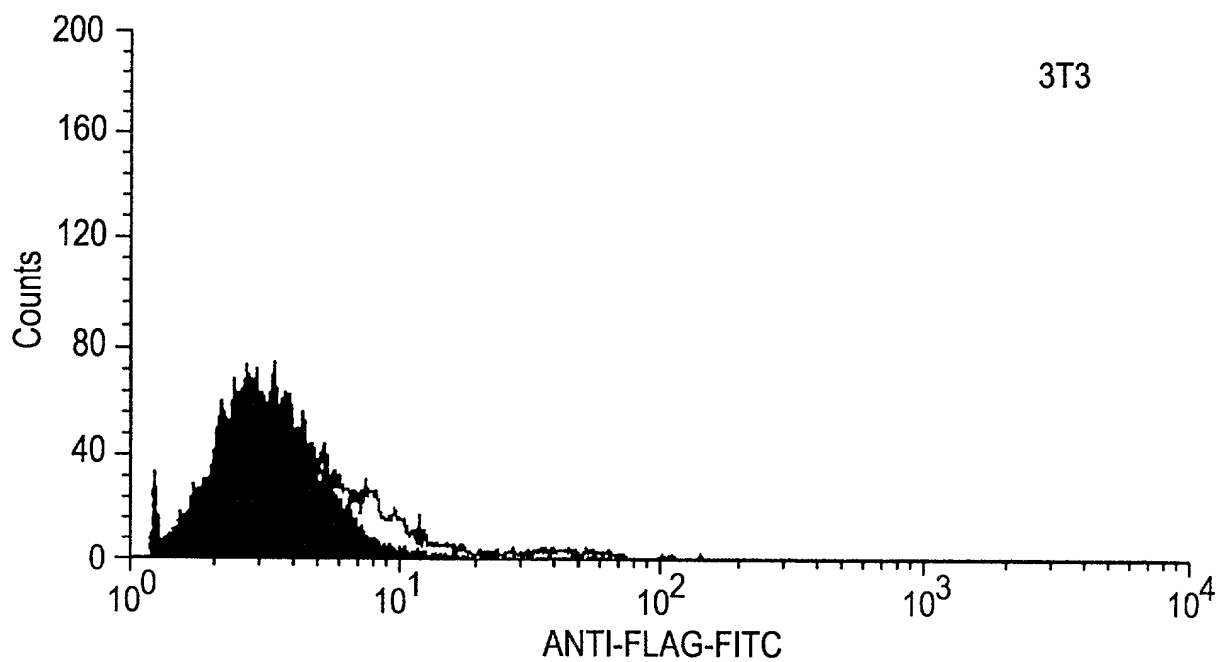
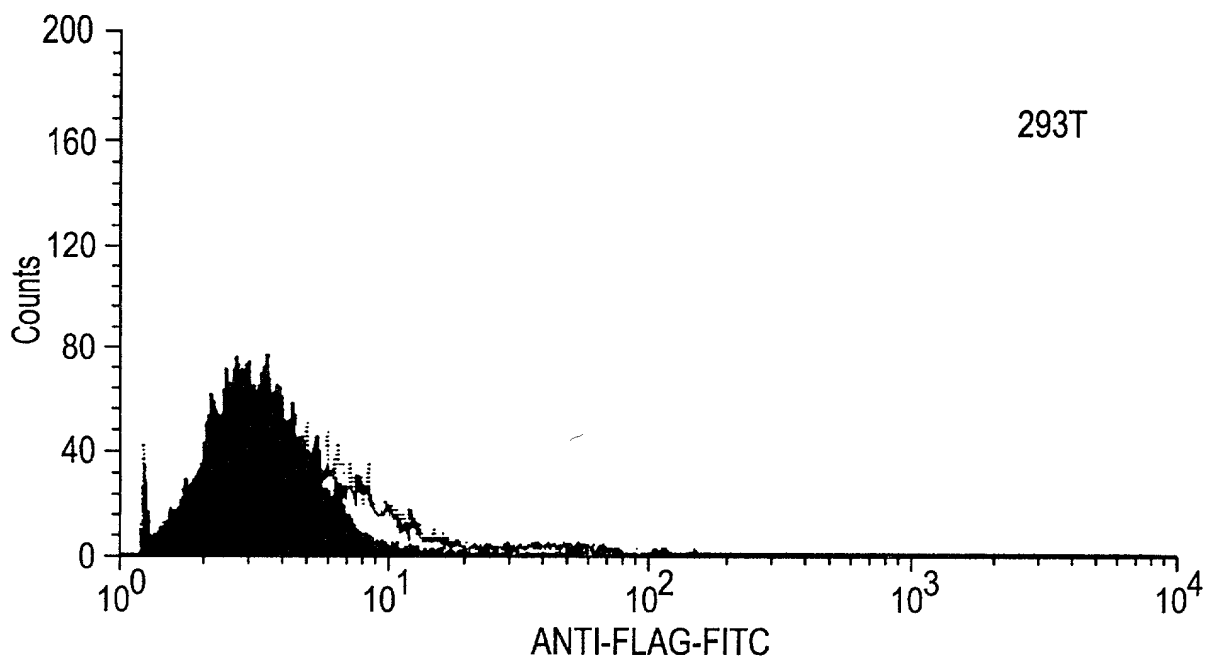


FIG. 5B-2



"09/854,864" 4984864

FIG. 5B-3

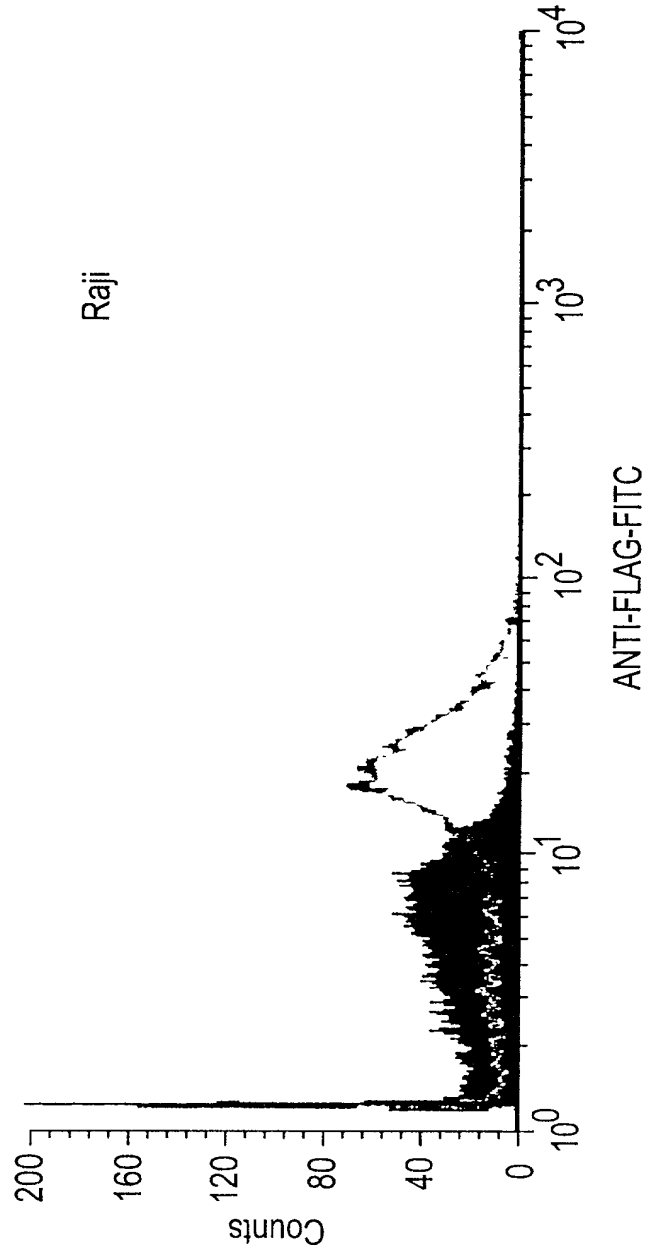
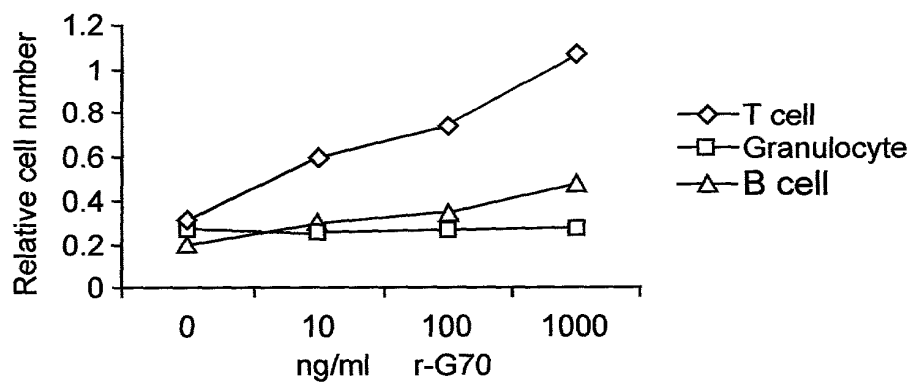


FIG. 6

The effect of r-G70/April on human
peripheral blood B cell, T cell and Granulocyte



The effect of IL-2 and G70/April on human
peripheral T cell proliferation

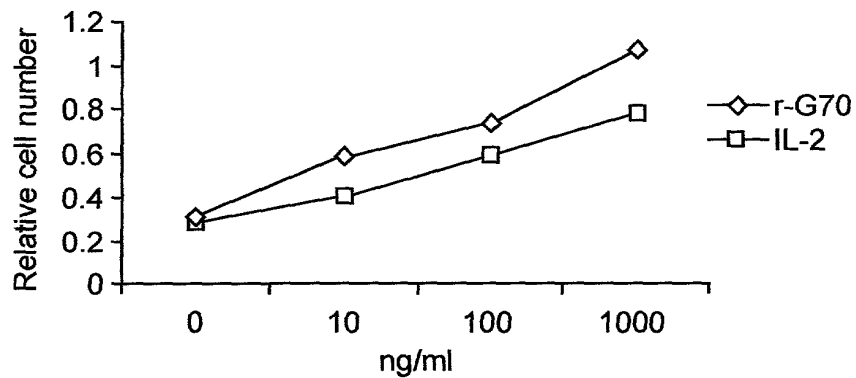
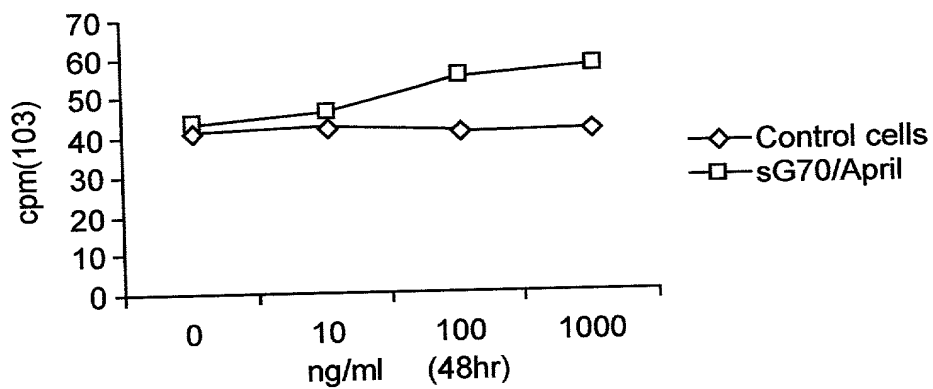


FIG. 7

Effect of sG70/April on murine B cell proliferation



Effect of sG70/April on murine T cell proliferation

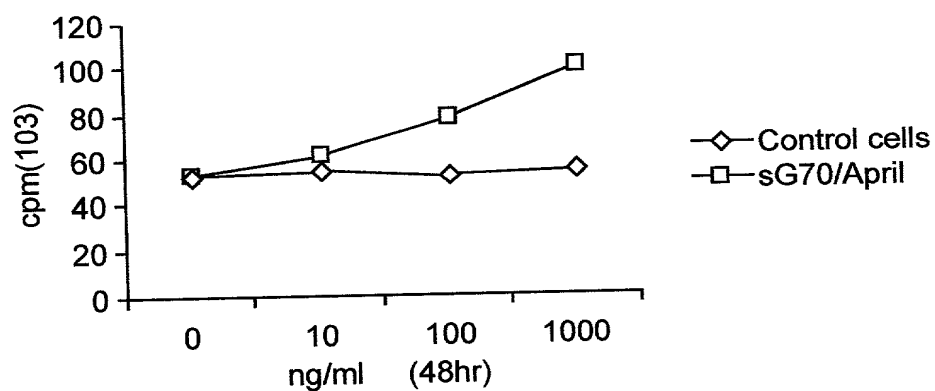


FIG. 8

Effect of G70/April on murine T cell
proliferation costimulated through CD28
antibody

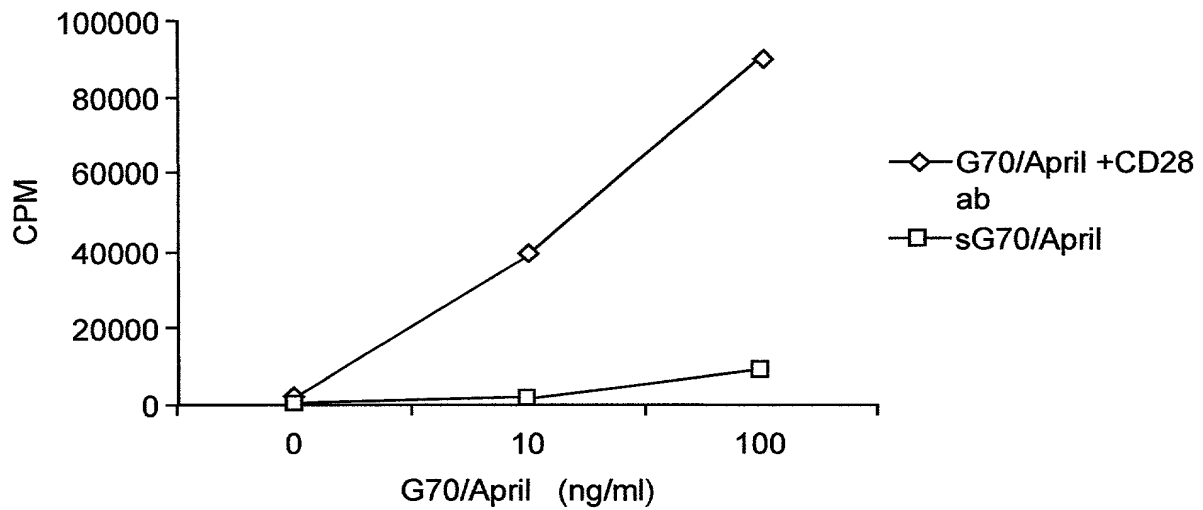


FIG. 9

Co-stimulatory activity of G70/April on mouse T cells

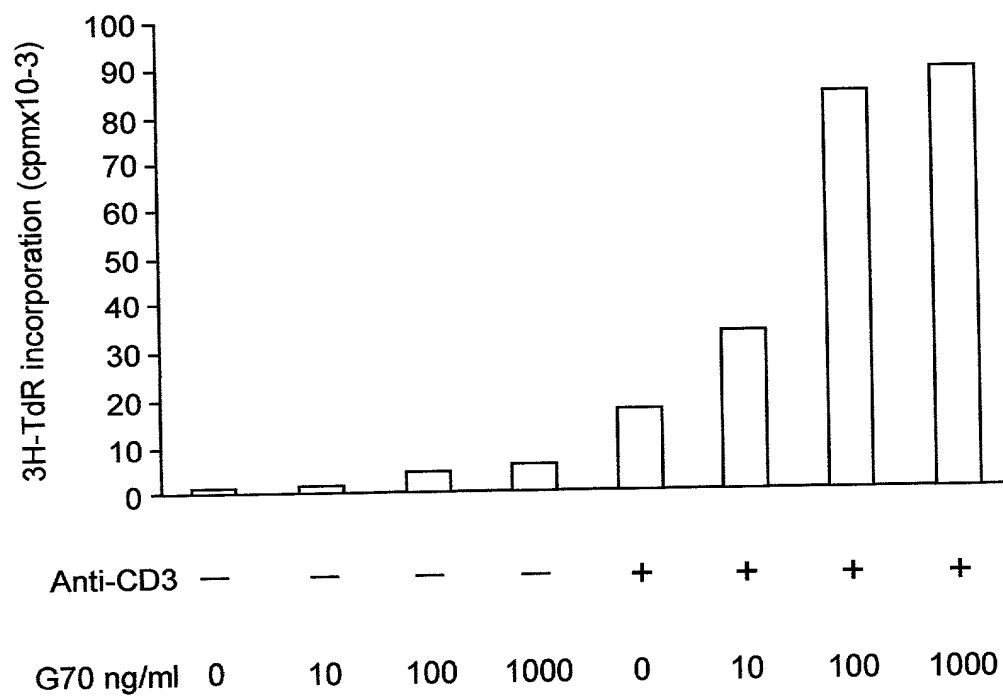


FIG. 10A

Human BCMA

Human (SEQ ID NO: 5):

1 MAGQCSQNEY FDSLLHACIP CQLRCSSNTP PLTCQRYCNA SVTNSVKGTN
51 AILWTCLGLS LIISLAVFVL MFLLRKISSE PLKDEFKNTG SGLLGMANID
101 LEKSRTGDEI ILPRGLEYTV EECTCEDCIK SKPKVSDHC FPLPAMEEGA
151 TILVTTKTND YCKSLPAALS ATEIEKSISA R

Human (SEQ ID NO: 5):

MAGQCSQ **NEYFDSLLHA CIPCQLRCSS NTPPLTCQRY** CNASVTNSVK
GTNA ILWTCL GLSLIISLAV FVLMFLLRKI SSEPLKDEFK NTGSGLLGMA
NIDLEKSRTG DEIILPRGLE YTVEECTCED CIKSKPKVDS DHCFLPAME
EGATILVTTK TNDYCKSLPA ALSATEIEKS ISAR

hBCMA's extracellular domain (SEQ ID NO: 6):

MAGQCSQ NEYFDSLLHA CIPCQLRCSS NTPPLTCQRY CNASVTNSVK
GTNA

hBCMA's cysteine-rich consensus region (SEQ ID NO: 7):

CSQ NEYFDSLLHA CIPCQLRCSS NTPPLTCQRY C

hBCMA's transmembrane region (SEQ ID NO: 8):

ILWTCL GLSLIISLAV FVLMF

FIG. 10B

huBCMA-Fc (SEQ ID NO: 9):

MAGQCSQNEYFDSLLHACIPCQLRCSSNTPPLTCQRYCNASVTNSVKGTNAGGG
GGDKTHTCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVK
FNWYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKAL
PAPIEKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNG
QPENNYKTTTPVLDSDGSFFLYSKLTVDKSRWQQGNVFSVMSVMHEALHNHYTQKS
LSLSPGK*

muBCMA-Fc (SEQ ID NO: 10):

MAQQCFHSEYFDSLLHACKPCHLRCSNPPATCQPYCDPSVTSSVKGSYTGGGGG
DKTHTCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFN
WYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPA
PIEKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQP
ENNYKTTTPVLDSDGSFFLYSKLTVDKSRWQQGNVFSVMSVMHEALHNHYTQKSLS
LSPGK*

FIG. 11

Alignment of human BCMA amino acid sequence and murine BCMA amino acid sequence

murine BCMA amino acid sequence Length: 185 (SEQ ID NO: 11):

```
1  MAQQCFHSEY FDSLLHACKP CHLRCSNPPA TCQPYCDPSV TSSVKGYTYV
51  LWIFLGLTLV LSLALFTISF LLRKMNPEAL KDEPQSPGQL DGSQALDKAD
101 TELTRIRAGD DRIFPRSLEY TVEECTCEDC VKSKPKGDS D HFFPLPAMEE
151 GATILVTTKT GDYKSSVPT ALQSVMGMEK PTHTR
```

alignment of human BCMA amino acid sequence and murine BCMA amino acid sequence.

```
Query:      4  MAGQCSQNEYFDSLLHACIPQCQLRCSNTPPLTCQRYCNASVTNSVKGTNAILWTCGLS 63
              MA QC  +EYFDSLLHAC PC LRCS+  PP TCQ YC+ SVT+SVKGT  +LW  LGL+
Sbjct:      1  MAQQCFHSEYFDSLLHACKPKCHLRCSN--PPATCQPYCDPSVTSSVKGYTYVWIFLGLT 58

Query:     64  LIISLAVFVLMFLLRKISSEPLKDEFKNTG----SGLLGMANIDLEKSRGTGDEIILPRGL 119
              L++SLA+F + FLLRK++ E LKDE ++ G   S  L  A+ +L + R GD+ I PR L
Sbjct:     59  LVLSLALFTISFLLRKMNPEALKDEPQSPGQLDGSQALDKADTELTRIRAGDDRIFRSL 118

Query:    120  EYTVEECTCEDCIKSKPKVDSHDHCFPLPAMEEGATILVTTKTNDYCKS-LPAAL-SATEI 177
              EYTVEECTCEDC+KSKPK DSDH FPLPAMEEGATILVTTKT DY KS +P AL S  +
Sbjct:    119  EYTVEECTCEDCVKSKPKGDSDDHFFPLPAMEEGATILVTTKTGDYKSSVPTALQSVMG 178

Query:    178  EKSISAR 184
              EK      R
Sbjct:    179  EKPTHTR 185
```


FIG. 12A

Human TACI

huTACI (SEQ ID NO: 14).

1 MSGLGRSRRG GRSRVDQEER FPQGLWTGVA MRSCPEEQYW DPLLGTCMSC
51 KTICNHQSQR TCAAFCRSLS CRKEQGKFYD HLLRDCISCA SICGQHPKQC
101 AYFCENKLRS PVNLPPELRR QRSGEVENNS DNSGRYQGLE HRGSEASPAL
151 PGLKLSADQV ALVYSTLGLC LCAVLCCFLV AVACFLKKRG DPCSCQPRSR
201 PRQSPAKSSQ DHAMEAGSPV STSPEPVETC SFCFPECRAP TQESAVTPGT
251 PDPTCAGRWG CHTRTTVLQP CPHIPDSGLG IVCVPAQEGG PGA

MSGLGRSRRGGRSRVDQEERFPQGLWTGVAMRSCPEEQYWDPLLGTCMSC
KTICNHQSQR TCAAFCRSLS CRKEQGKFYD HLLRDCISCASICGQHPKQC
AYFCENKLRS PVNLPPELRR QRSGEVENNS DNSGRYQGLE HRGSEASPAL
PGLKLSADQV ALVYSTLGLC LCAVLCCFLV AVACFLKKRG DPCSCQPRSR
PRQSPAKSSQ DHAMEAGSPV STSPEPVETC SFCFPECRAP TQESAVTPGT
PDPTCAGRWG CHTRTTVLQP CPHIPDSGLG IVCVPAQEGG PGA

huTACI's extracellular domain (SEQ ID NO: 15):

1 MSGLGRSRRG GRSRVDQEER FPQGLWTGVA MRSCPEEQYW DPLLGTCMSC
51 KTICNHQSQR TCAAFCRSLS CRKEQGKFYD HLLRDCISCA SICGQHPKQC
101 AYFCENKLRS PVNLPPELRR QRSGEVENNS DNSGRYQGLE HRGSEASPAL
151 PGLKLSADQV ALVYST

FIG. 12B

huTACI's cysteine-rich consensus region (SEQ ID NO: 16):

CPEEQYWDPLLGTCTMSCKTICNHQSQR TCAAF C and
CRKEQGKFYDHLRLDCISCASICGQHPKQCAYFC

transmembrane region (SEQ ID NO: 17):

LGLCLCAVLCCFLVAVACFL

hTACI-Fc (SEQ ID NO: 18):

1 MSGLGRSRRG GRSRVDQEER FPQGLWTGVA MRSCPEEQYW DPLLGTCTMSC
51 KTICNHQSQR TCAAFCRSL S CRKEQGKFYD HLLRDCISCA SICGQHPKQC
101 AYFCENKLRS PVNLPPELRR QRSGEVENNS DNSGRYQGLE HRGSEASPAL
151 PGLKLSADQV ALVYSGGGGG DKTHTCPPCP APELLGGPSV FLFPKPKD
201 LMISRTPEVT CVVVDVSHED PEVKFNWYVD GVEVHNAKTK PREEQYNSTY
251 RVVSVLTVLH QDWLNGKEYK CKVSNKALPA PIEKTISKAK GQPREPQVYT
301 LPPSRDELTK NQVSLTCLVK GFYPSDIAVE WESNGQPENN YKTTTPVLDS
351 DGSFFLYSKL TVDKSRWQQG NVFSCSV MHE ALHNHYTQKS LSLSPGK*

FIG. 13

Alignment of cysteine rich extracellular regions of human TACI and human BCMA.

```
34 CPEEQYWDPLLGTCTMSCKTICNHQS.QRTCAAFCRSLSCRKEQGKFYDHL 82
   | : :|. | || |. |. |. . || :| . . | . :
8  CSQNEYFDSLLHACIPCQLRCSSNTPPLTCQRYCNASVTNSVKGT..NAI 55

      .
83 LRDCISCASI 92
   | | : . |
56 LWTCLGLSLI 65
```

FIG. 14A

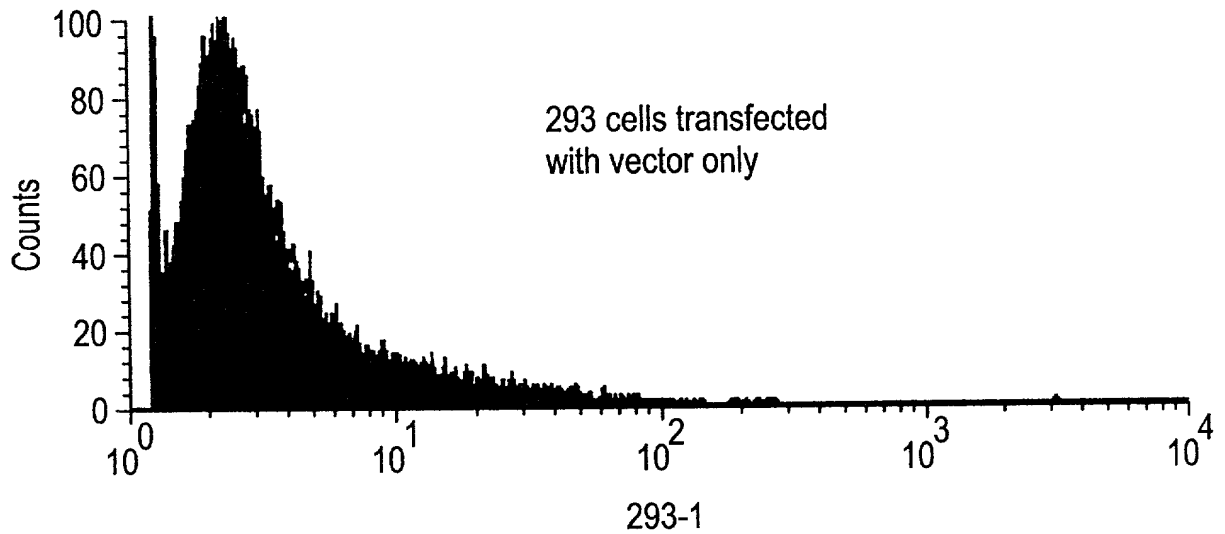


FIG. 14B

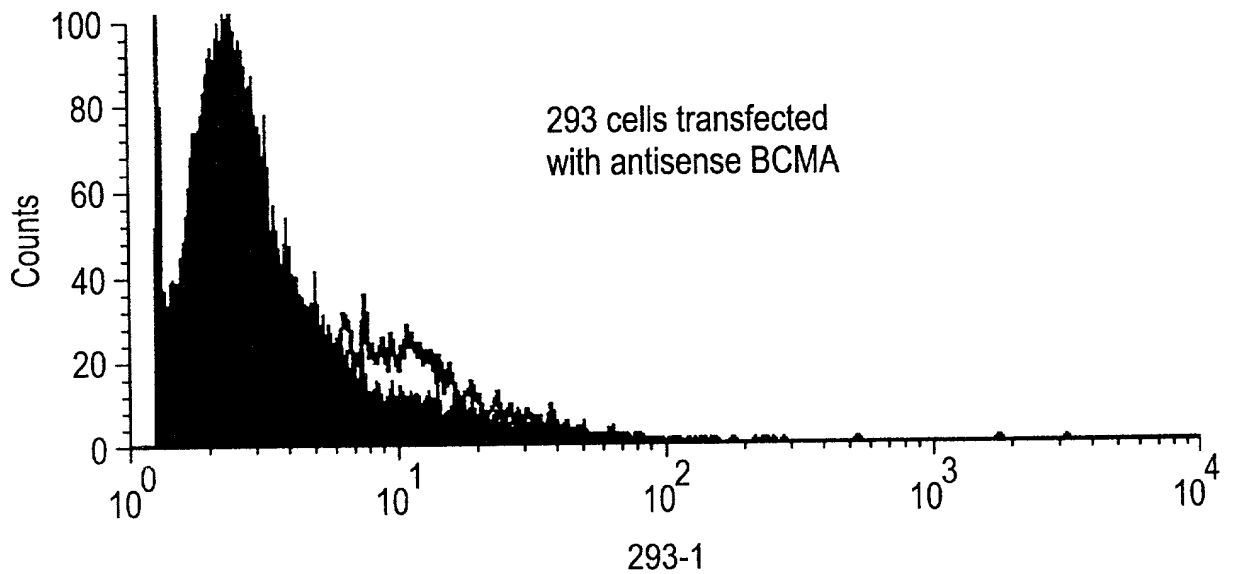


FIG. 14C

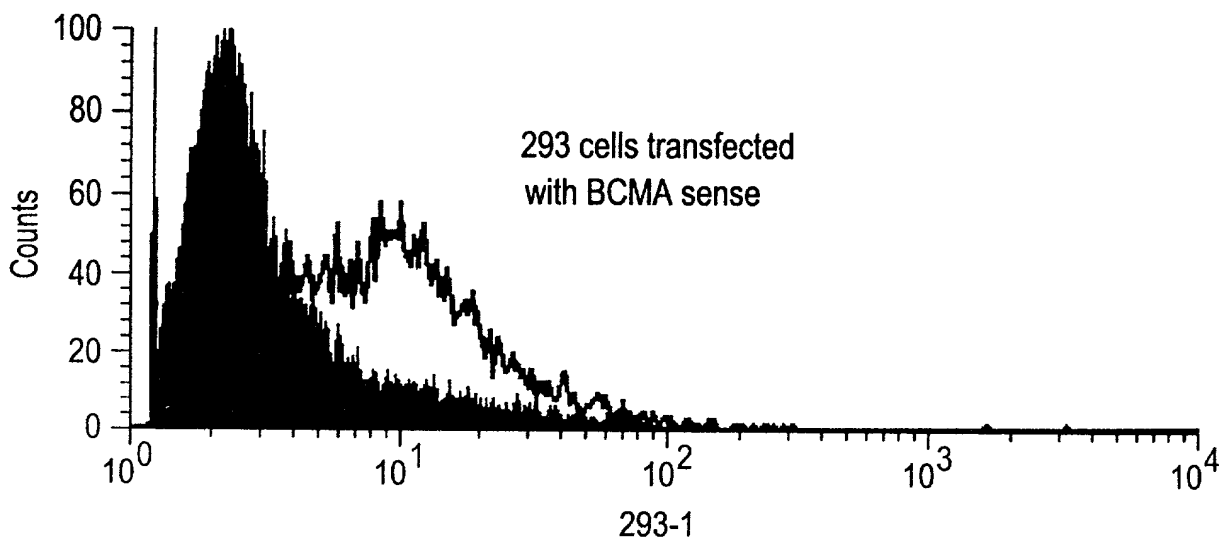


FIG. 15A

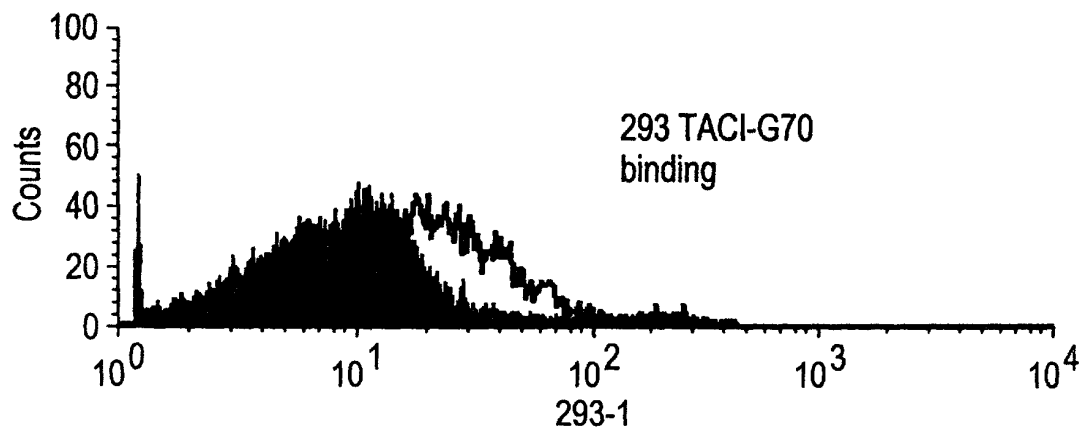


FIG. 15B

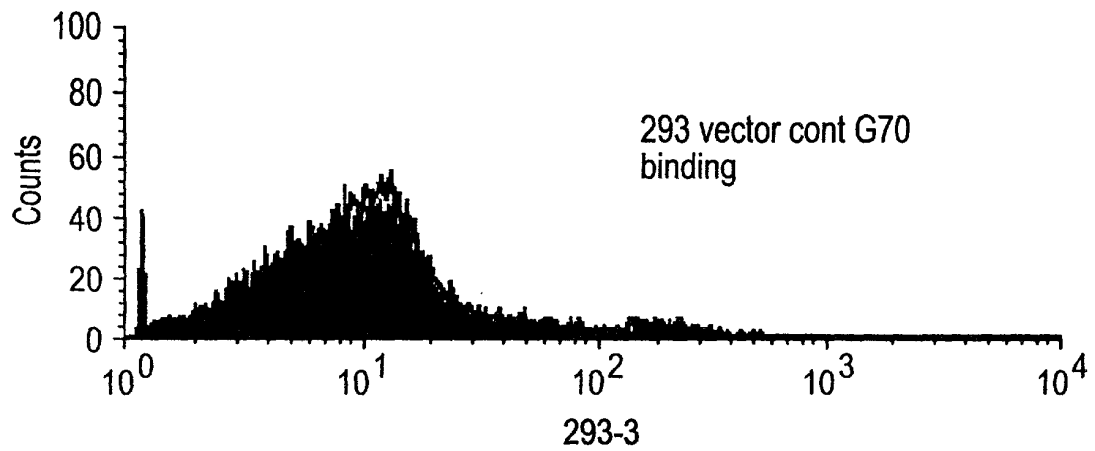


FIG. 16A

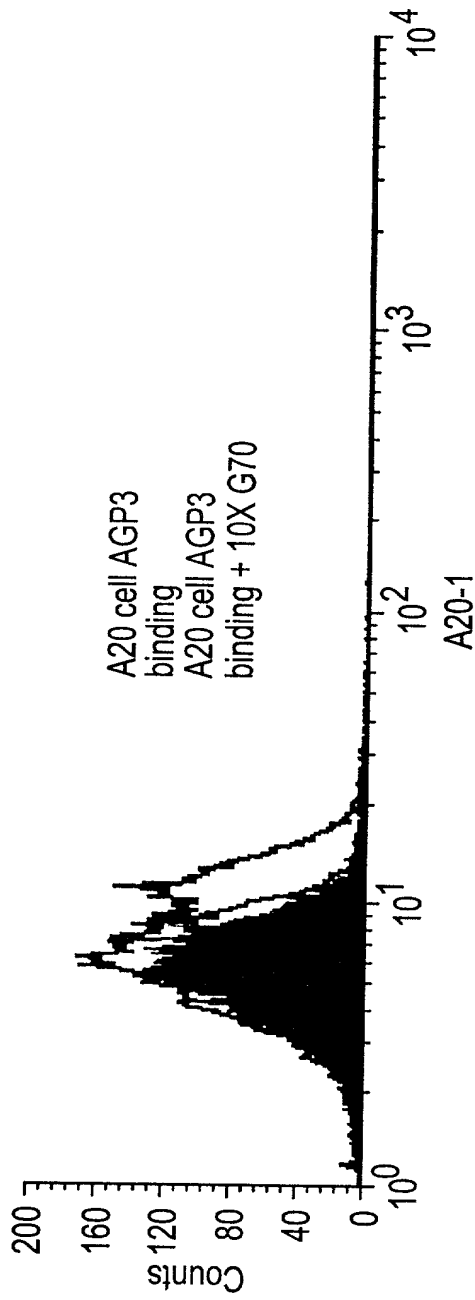
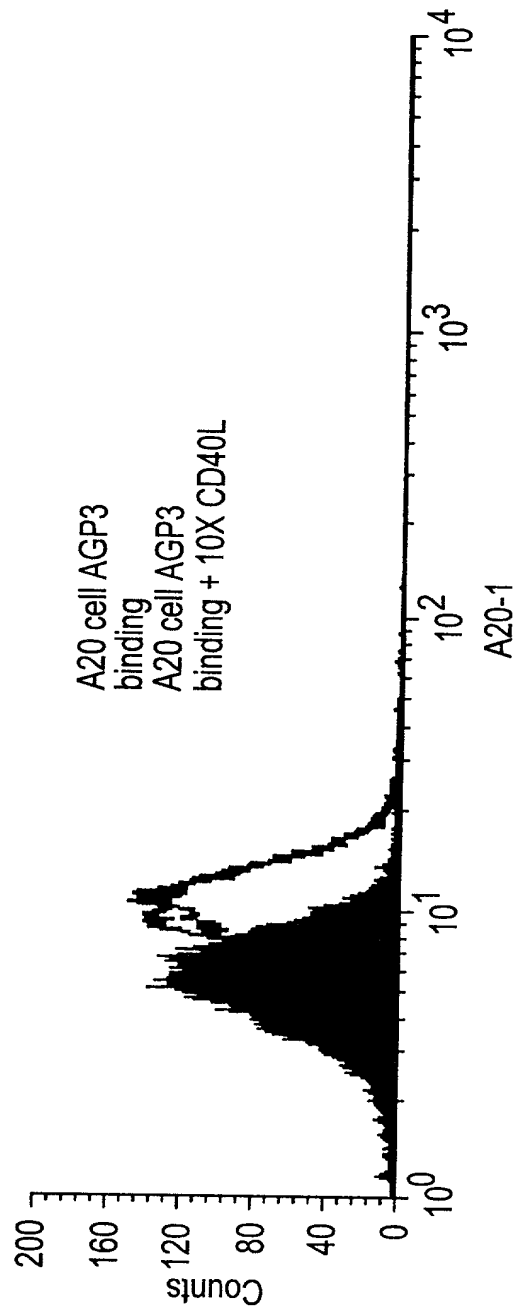


FIG. 16B



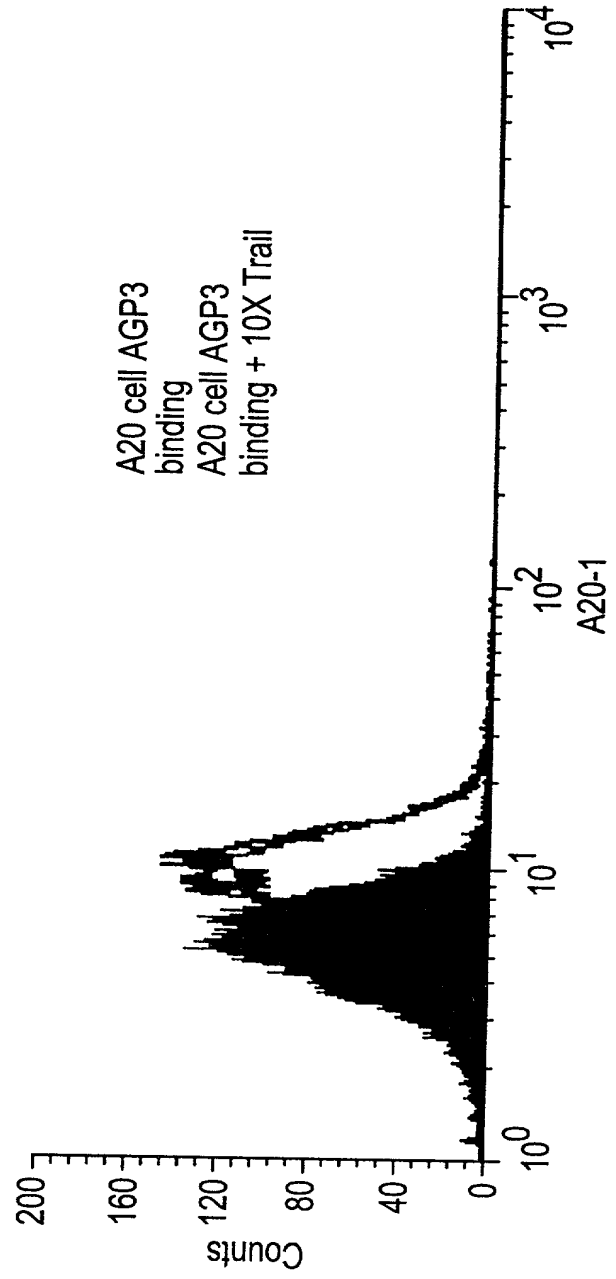
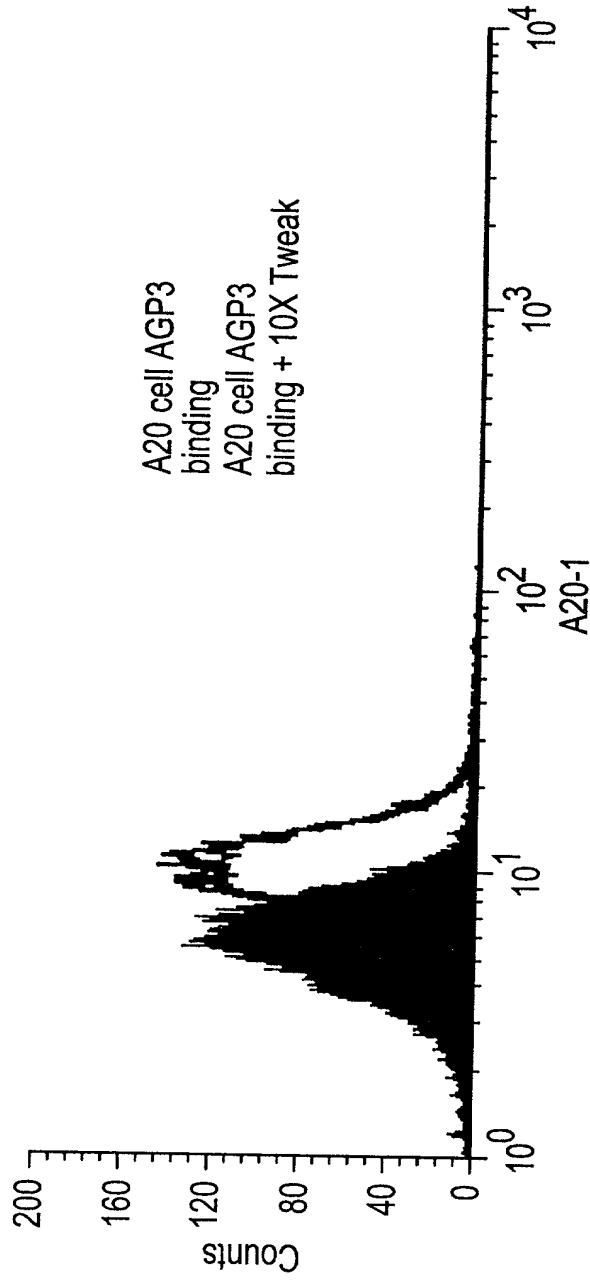


FIG. 16C

FIG. 16D

FIG. 16C

FIG. 17A

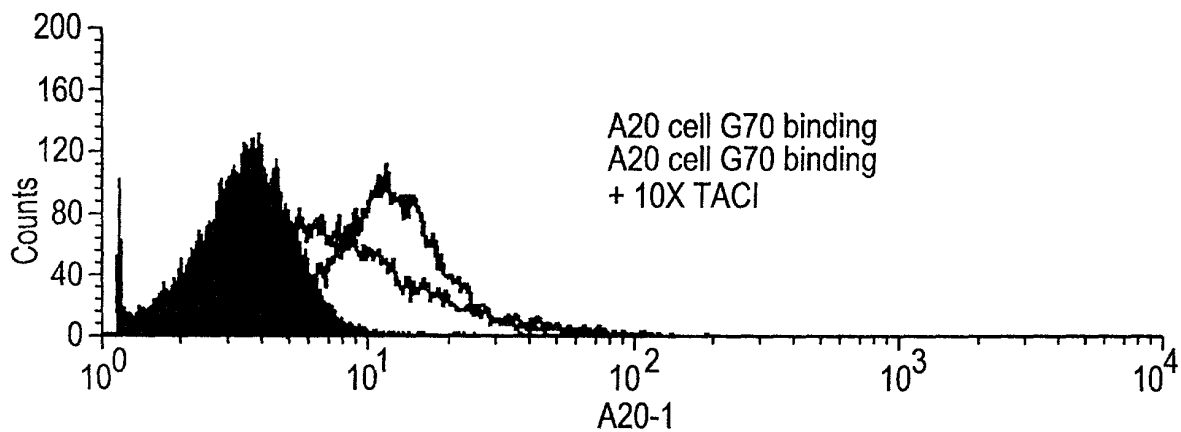


FIG. 17B

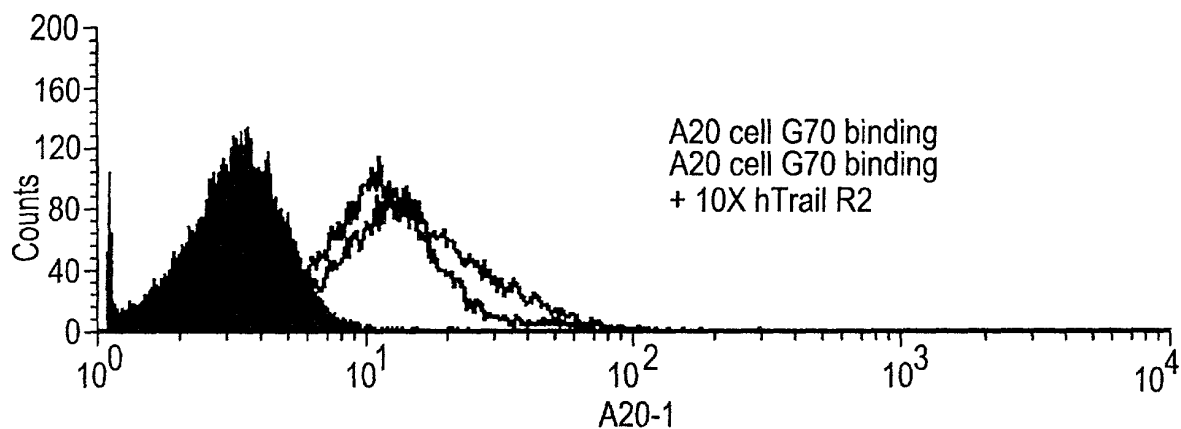


FIG. 17C

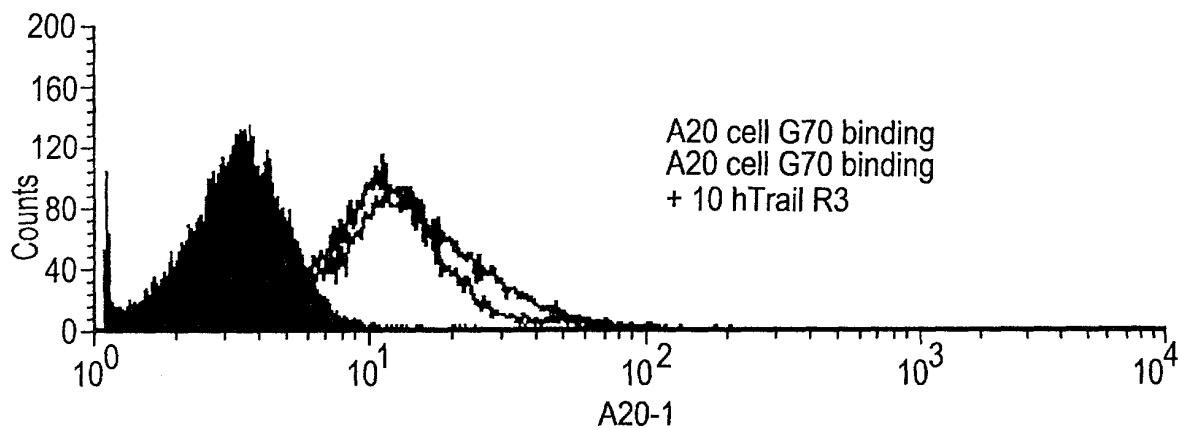


FIG. 18

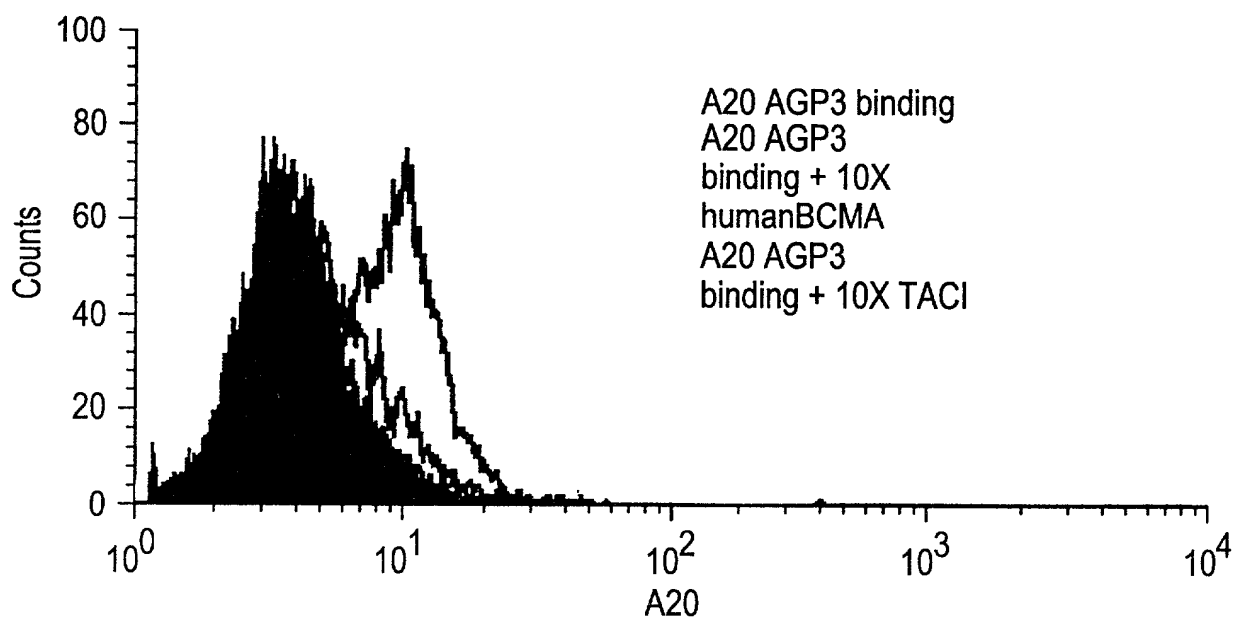


FIG. 19A

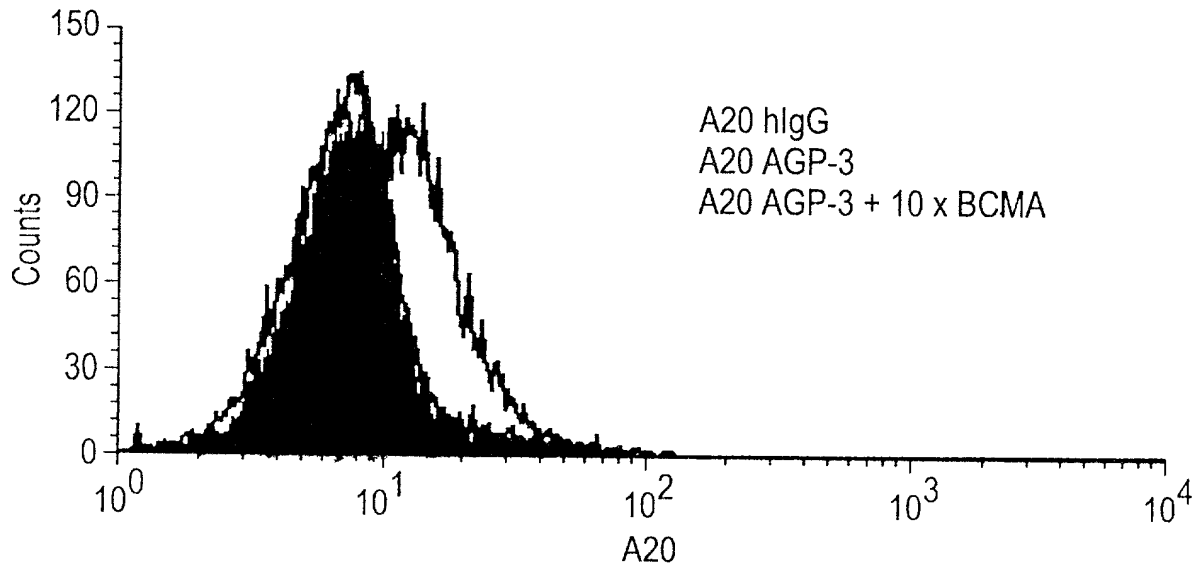


FIG. 19B

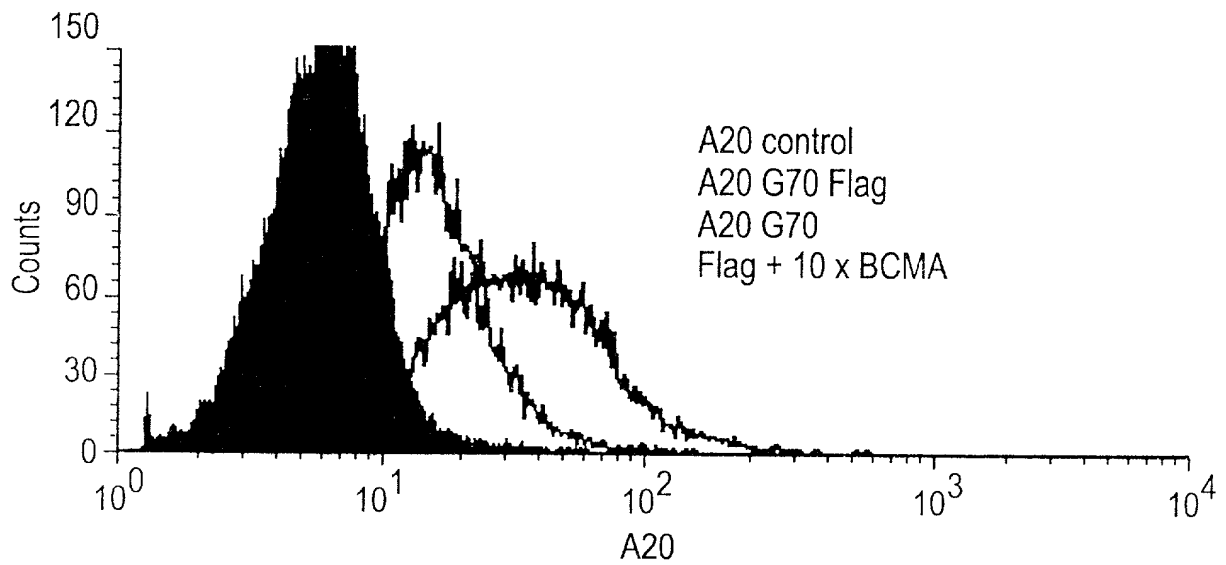


FIG. 20A

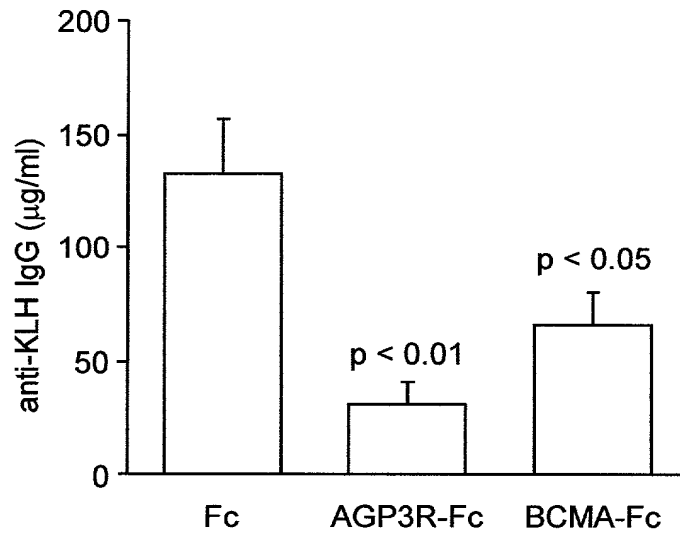


FIG. 20B

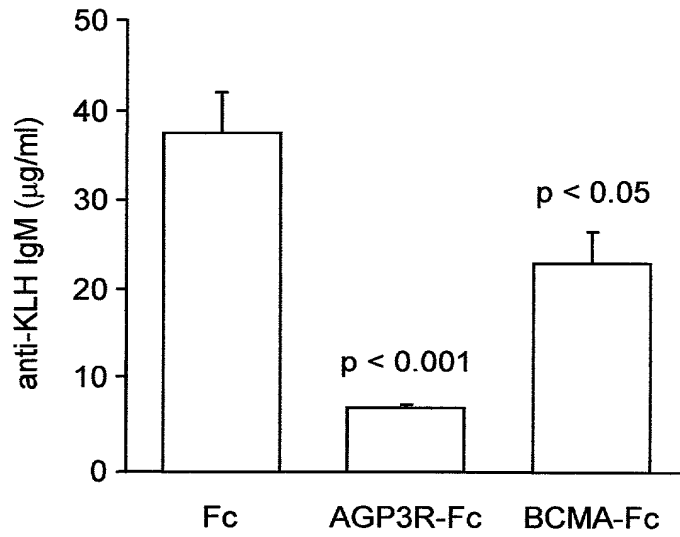
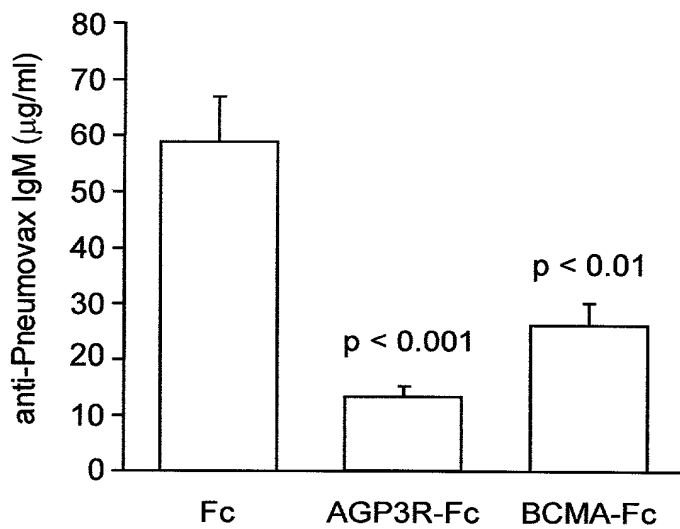


FIG. 20C



"4346" 494569

FIG. 21

Fc-humanAPRIL

Fc-humanAPRIL protein sequence including the signal sequence, Fc domain, linker (Xhol site) and APRIL:

1	MEWSWVFLFF	LSVTTGVS	HD	KTH	TCPPCPA	PELLGGPSVF
	LFPPKPKDTL					
51	MISRTPEVTC	VVVDVSHEDP	EVKFNWYVDG	VEVHNAKTKP		
	REEQYNSTYR					
101	VVSVLTVLHQ	DWLNGKEYKC	KVSNKALPAP	IEKTISKAKG		
	QPREPQVYTL					
151	PPSRDELTKN	QVSLTCLVKG	FYPSDIAVEW	ESNGQPENNY		
	KTTTPPVLDSD					
201	GSFFLYSKLT	VDKSRWQQGN	VFSCSVMHEA	LHNHYTQKSL		
	SLSPGK	SRAV				
251	LTQKQKKQHS	VLHLVPINAT	SKDDSDVTEV	MWQPALRRGR		
	GLQAQGYGVR					
301	IQDAGVYLLY	SQVLFQDVTF	TMGQVVSREG	QGRQETLFR		
	IRSMPSHPDR					
351	AYNSCYSAGV	FHLHQGDILS	VIIPRARAKL	NLSPHGFTFLG		
	FVKL*					

FIG. 22
Fc-HumanAPRIL and soluble human AGP3
stimulate proliferation of primary B cells

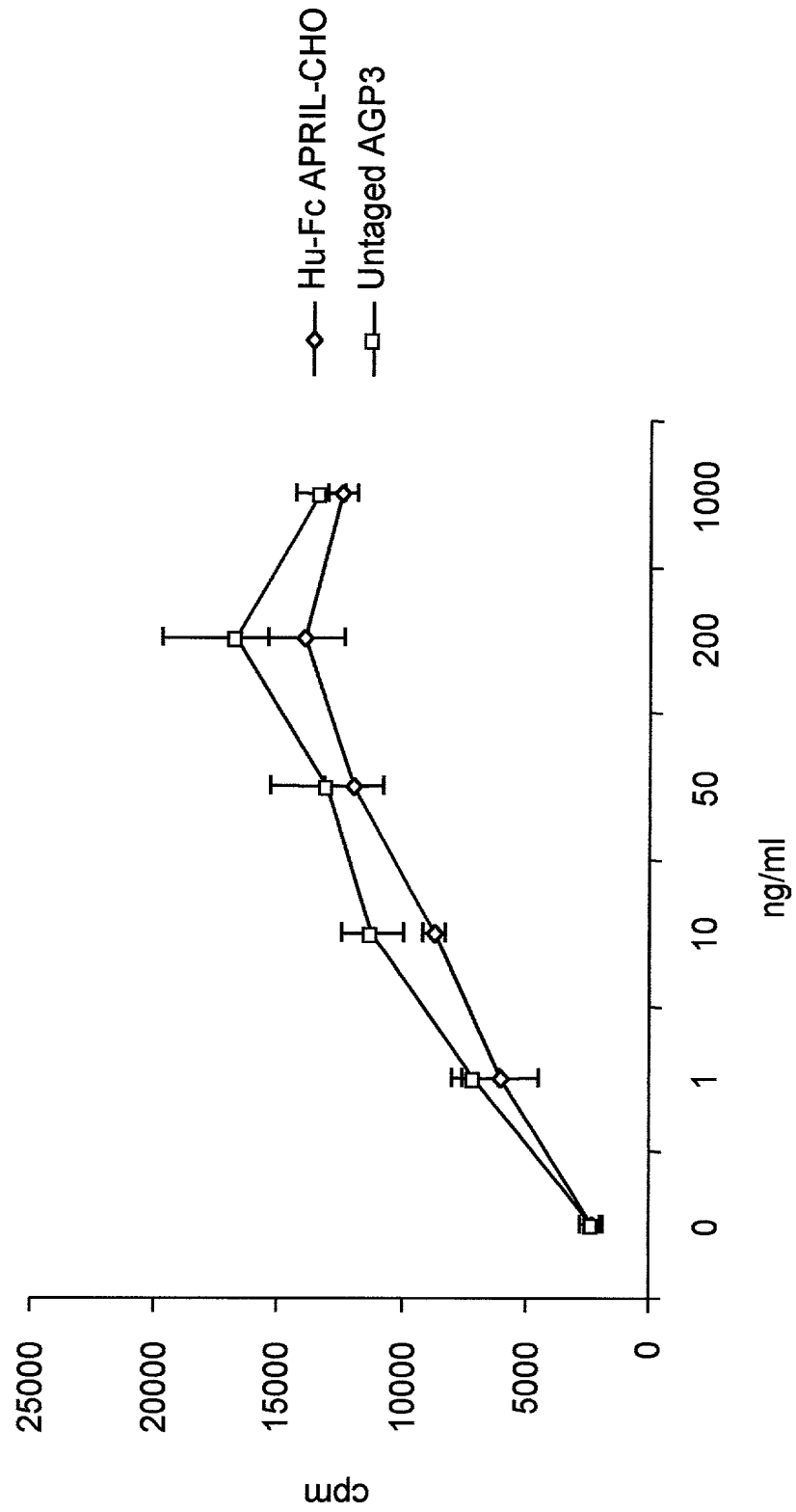
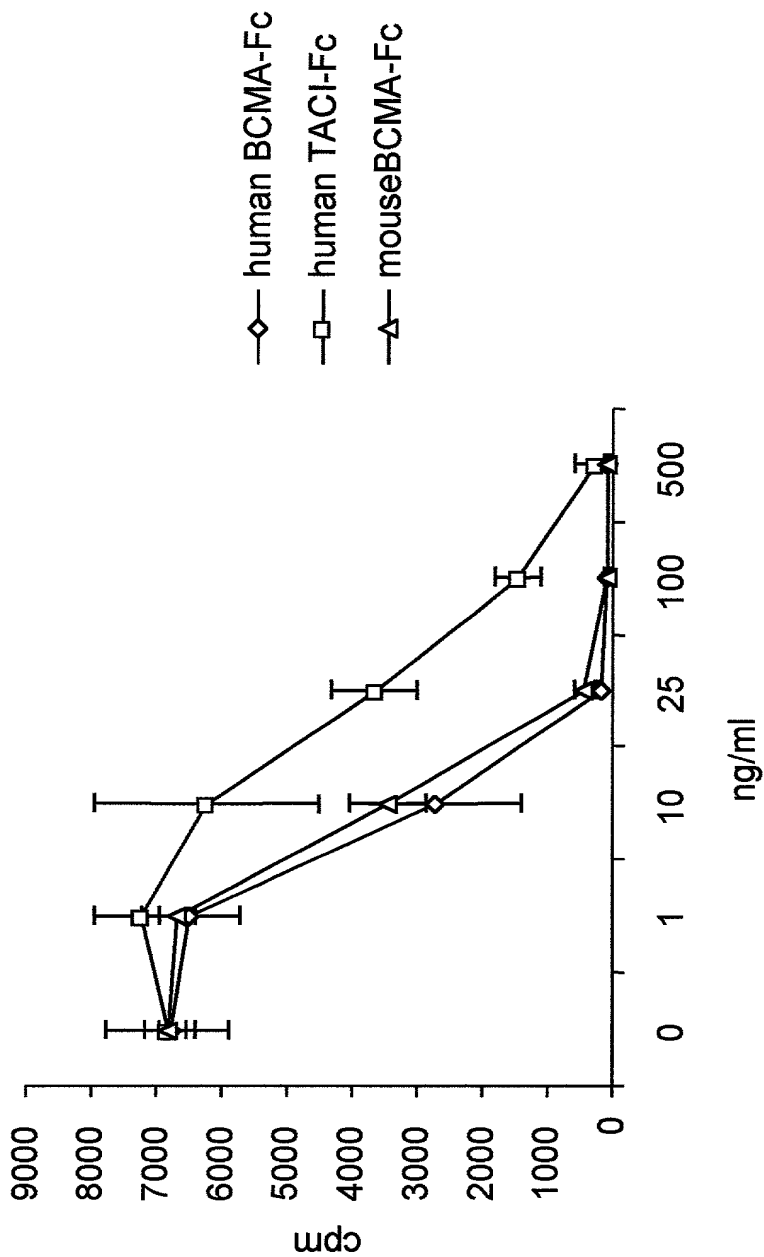


FIG. 23

hBCMA-Fc and wt hTACI-Fc inhibits
Flag-mAPRIL mediated mouse B cell
proliferation



49/854,864 49/854,864

FIG. 24
hBCMA-Fc reduces PB B cell level *in vivo*
15 mg/kg ip on day 0, 3, and 6

BLOOD		WBC 10e6/ml	#Lym 10e6/ml	CD3+ #	CD3-B220+ #
BCMA-Fc SD t test		5.30 0.39 0.03318	3.81 0.43 0.01570	2.3 0.32 0.24737	1.3 0.27 0.00506
		8.02 1.27	6.43 1.52	2.7 0.6	3.2 0.6
FC SD		6.90 2.04	5.55 1.79	2.1 0.5	2.9 1.2

FIG. 25

hBCMA-Fc reduces spleen B cell levels *in vivo*
15 mg/kg ip on day 0, 3, and 6

Spleen	WBC 10e6/ml	Lym (%)	spleen lym# 10ml(x10e6)	CD3-B220+ (%)	CD3-B220+ #
BCMA-Fc SD t test	9.12 0.92 0.02778	97.9 0.51 0.89118	89.3 9.32 0.02668	45.5 1.29 0.00234	41.8 4.92 0.02088
Fc SD	11.49 1.62	97.9 0.38	112.5 15.65	50.6 1.95	57.1 9.67
Saline SD	11.48 1.71	98.5 0.1	113.1 16.9	53.7 6.7	48.5 29.15

FIG. 26

Flag-mAPRIL and hAGP3 mediated IgA production
 inhibited by hBCMA-Fc and hTACI-Fc *in vitro*

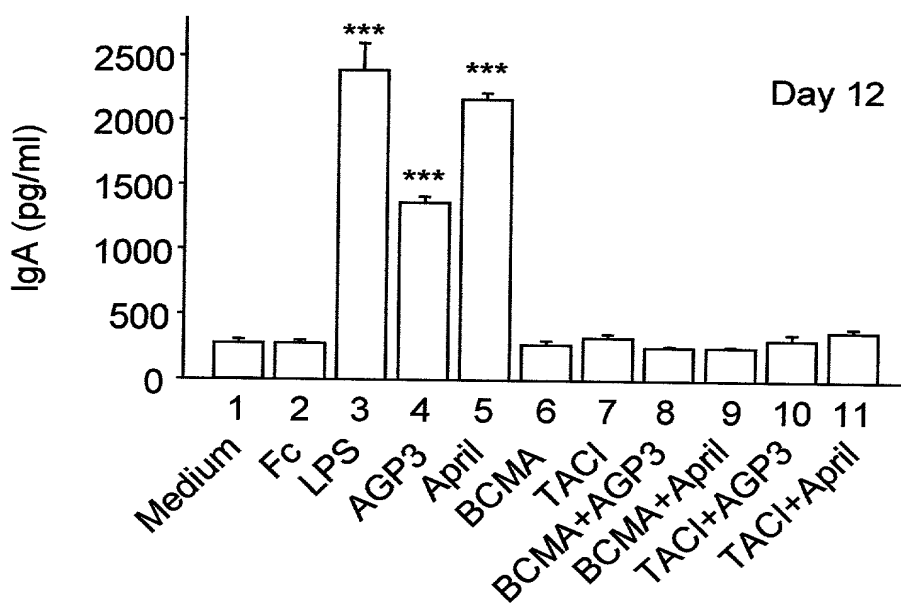
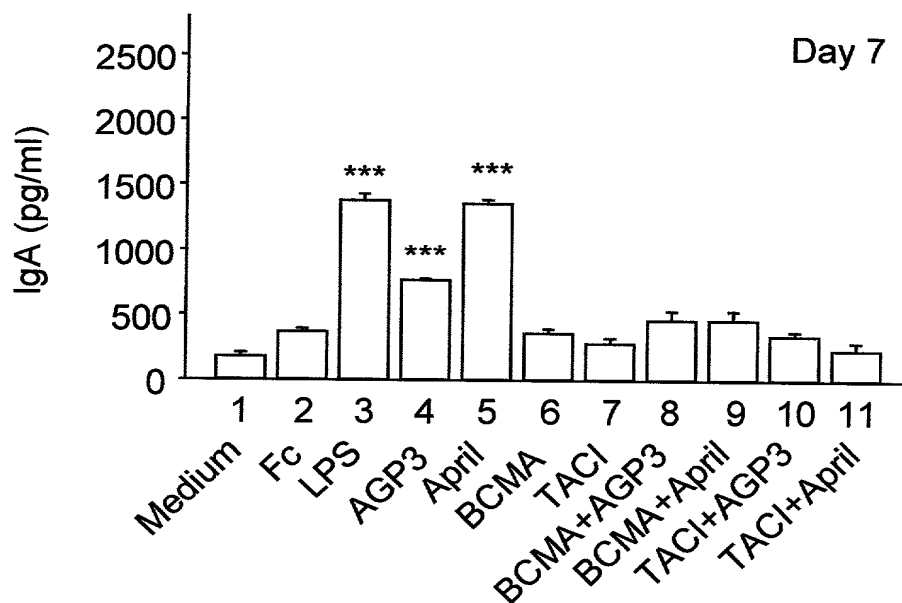


FIG. 27

Flag-mAPRIL and hAGP3 Mediated IgG Production
 Inhibited by BCMA-Fc and TACI-Fc *in Vitro*

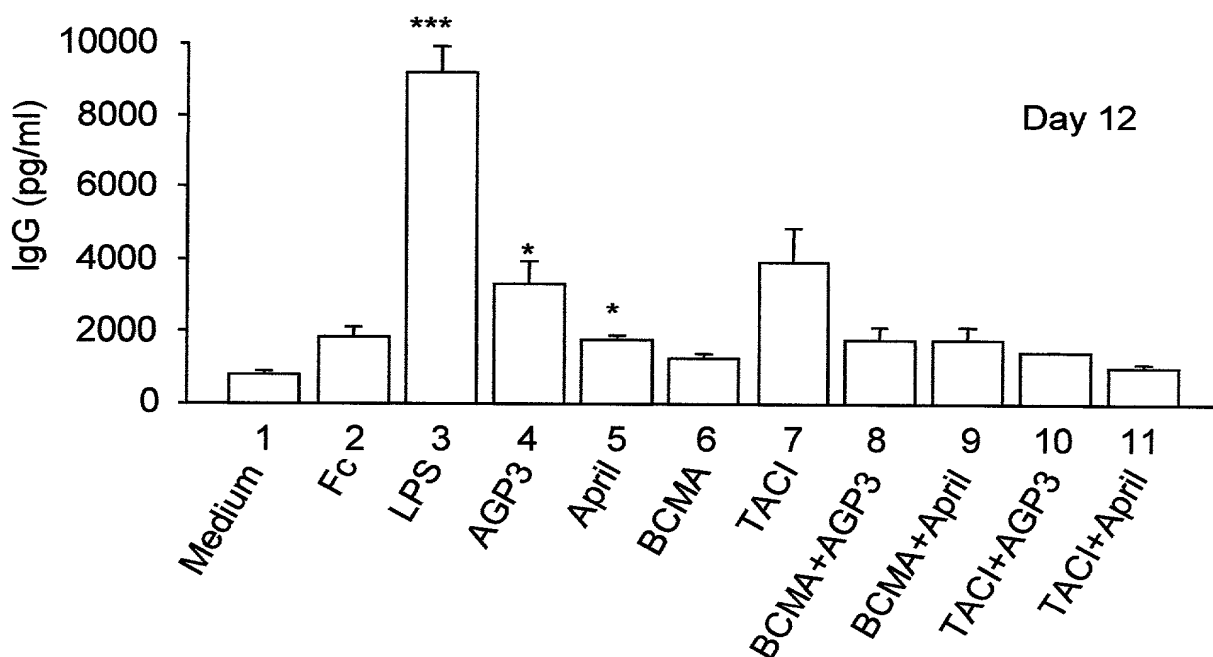
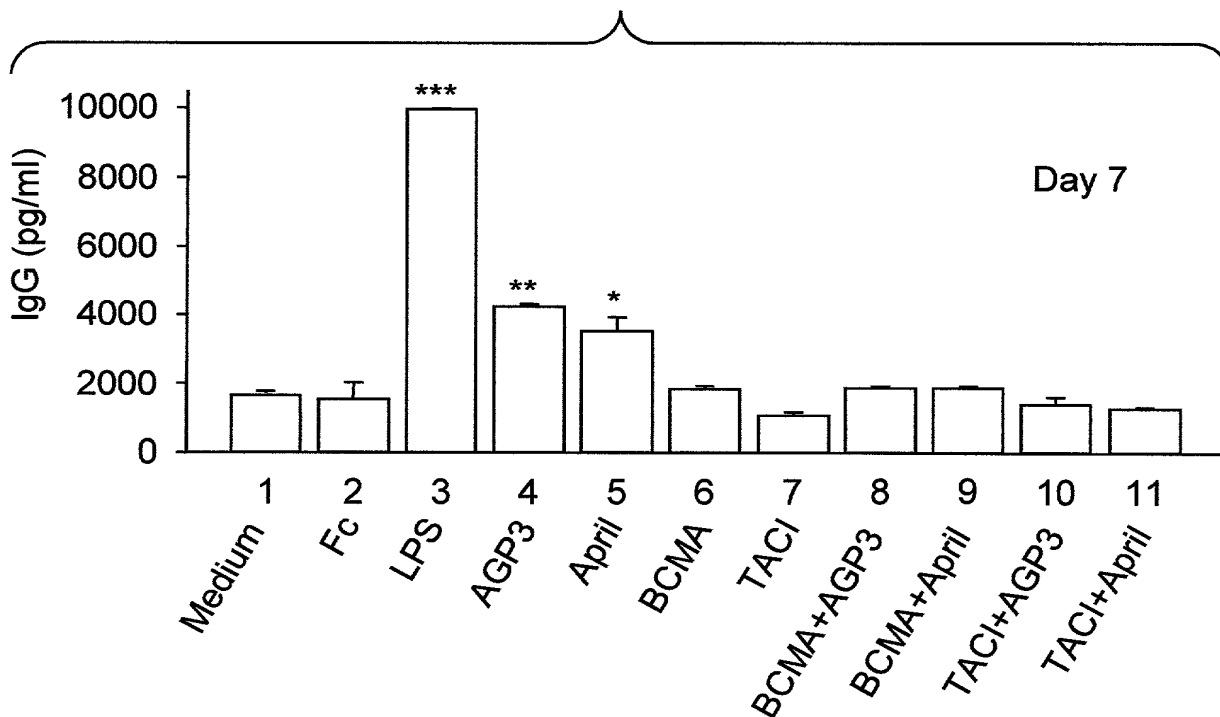


FIG. 28

Significantly reduces total IgE and IgA in normal mice treated with mBCMA-Fc and trun hTACI-Fc 5 mg/kg ip day 0, 3, and 6

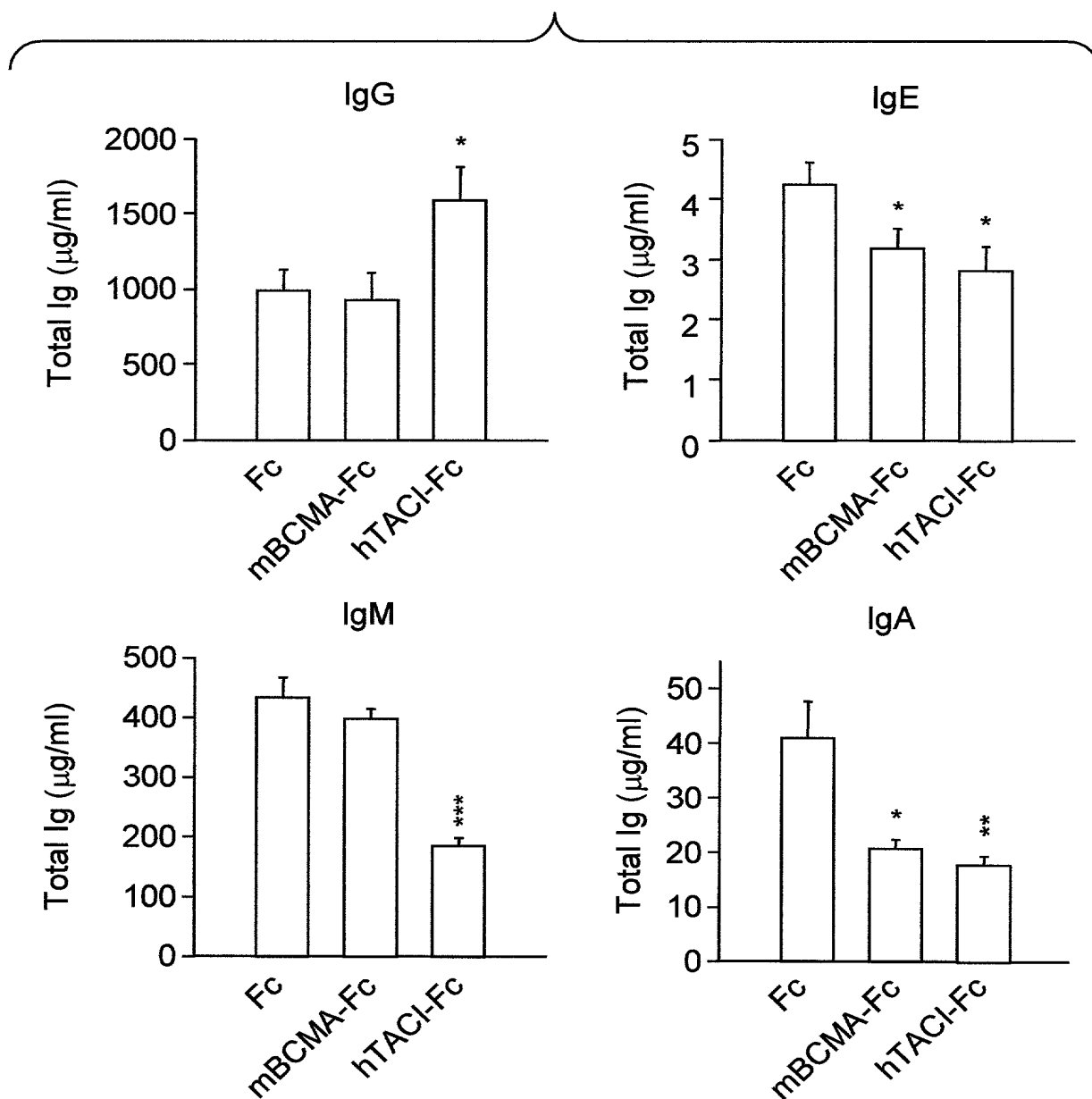


FIG. 29

BCMA-Fc and truncated TACI-Fc at daily doses of 0.5 mg/kg inhibits humoral immunity *in vivo*

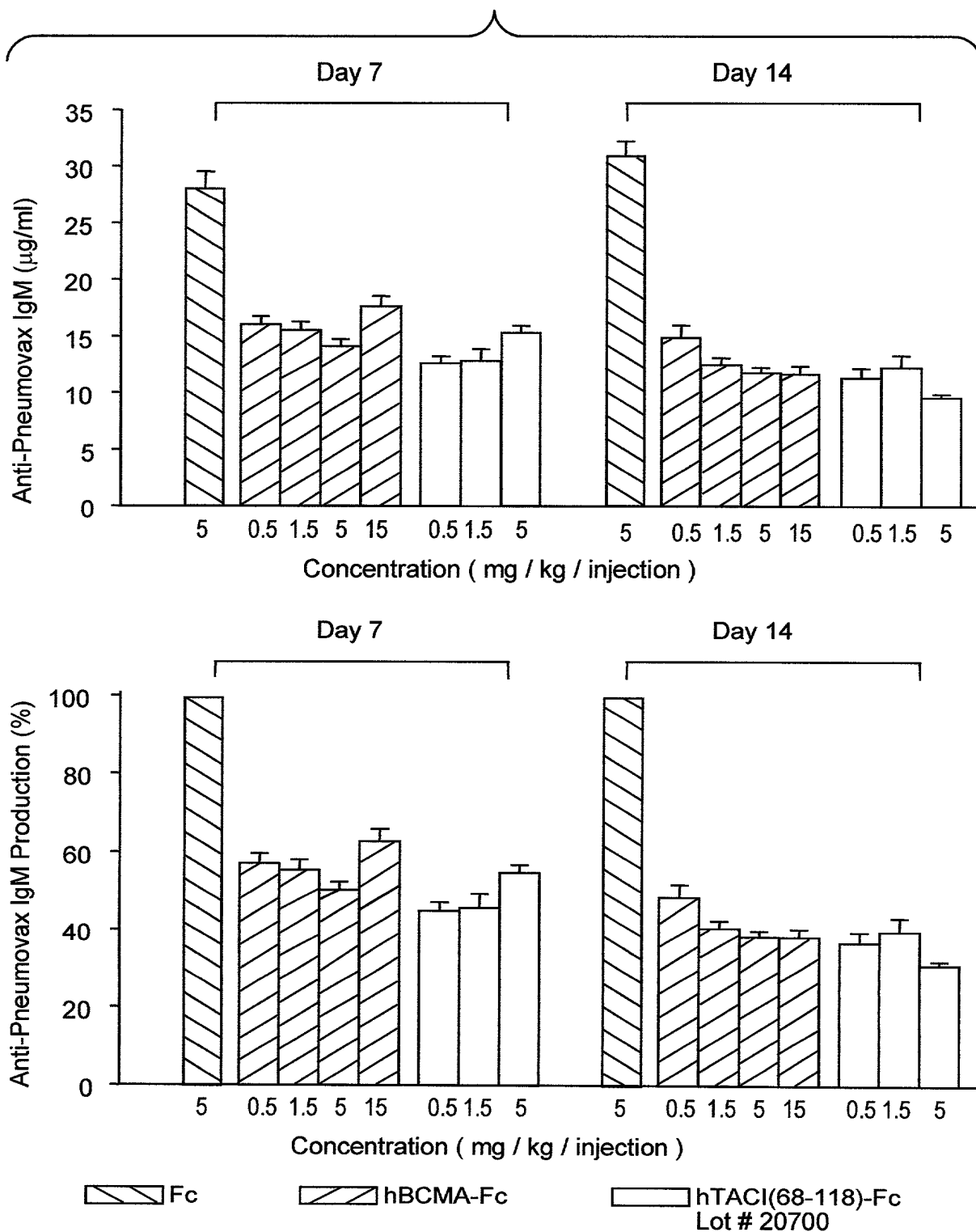


FIG. 30

Anti-mAPRIL c-19 MAb
Inhibition of APRIL mediated B cell proliferation

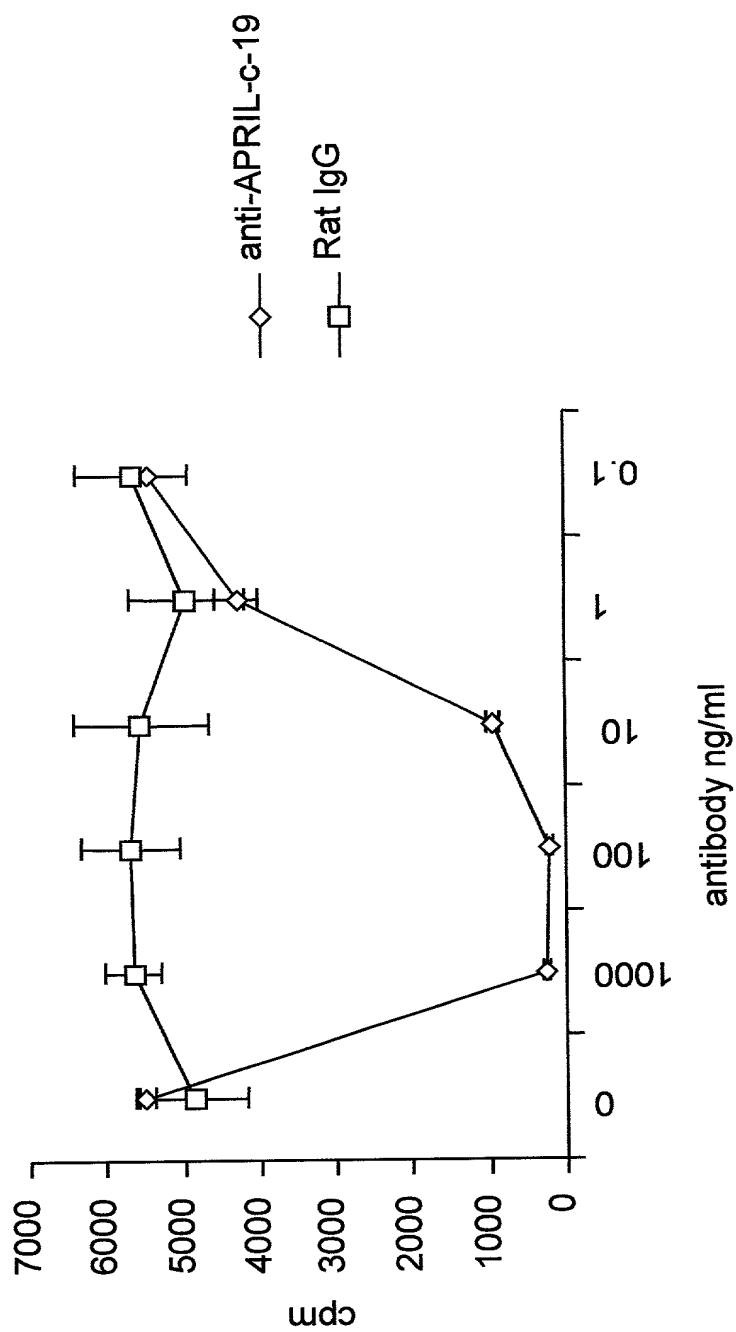
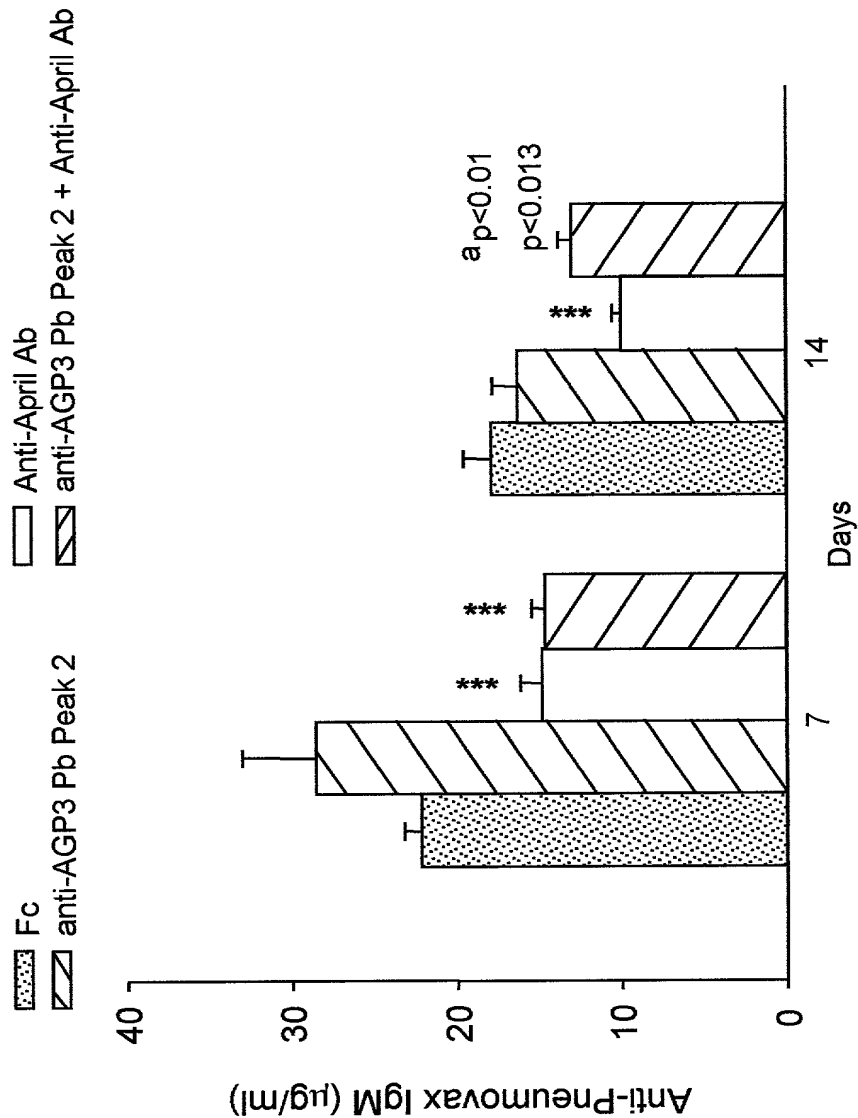
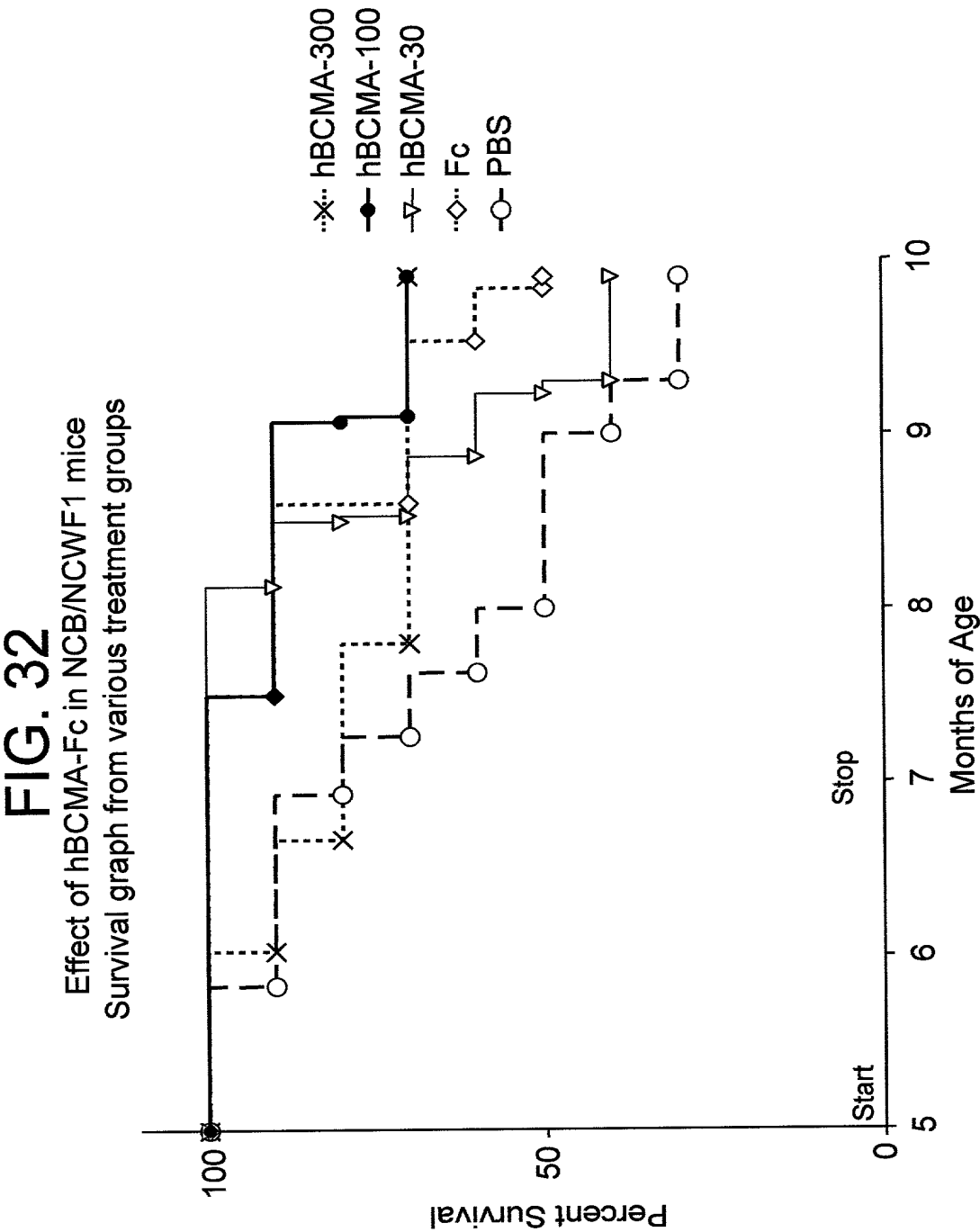


FIG. 31
Neutralizing anti-mAPRIL Mab Reduces anti-Pneumovacs IgM *In Vivo*
5 mg/kg ip on day 0, 3, and 6

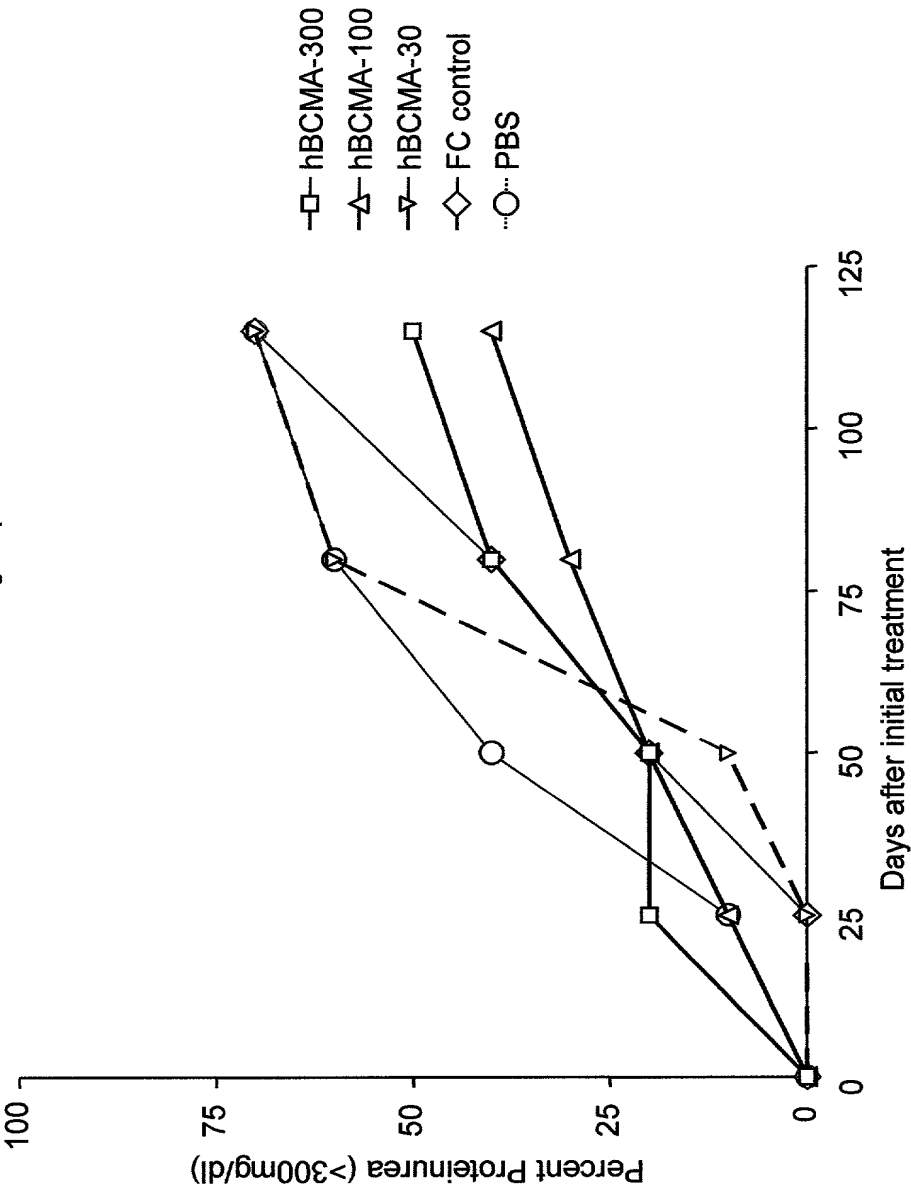


^a difference between Anti-APRIL Ab and anti-AGP3 Pb Peak 2 + Anti-APRIL Ab Groups



N=10 Mice were treated for 8 weeks 3x/week with the indicated proteins. KIN2 group had 12 mice.
The 100 in the legend stands for 100 µg of protein or 4mg/kg i.p.

FIG. 33
Effect of hBCMA-Fc in NCB/NCWF1 mice
Percentage of mice with proteinuria (>300mg/dl)
from various treatment groups



N=10 Five month old BWF1 mice were treated with protein for 8 weeks i.p.
The hBCMA-300 stands for hBCMA-fc 300 µg/mouse (12mg/kg)

FIG. 34

Analysis of antibodies to dsDNA from the peripheral blood
from various treatment groups of BWF1 at day 0,30,60, and 90.

MEAN anti-dsDNA isotypes in U/ml

Group #	Day 0		Day 30		Day 60		Day 90	
	IgG	IgM	IgG	IgM	IgG	IgM	IgG	IgM
hBCMA-300	179	560	163	371	150	706	171	841
hBCMA-100	150	430	259	718	171	822	339	1031
hBCMA-30	377	592	297	458	401	664	424	601
FC.	149	371	234	283	384	331	432	351
PBS	308	292	439	311	247	576	720	467

Standard Deviation of the above means

Group #	Day 0		Day 30		Day 60		Day 90	
	IgG	IgM	IgG	IgM	IgG	IgM	IgG	IgM
hBCMA-300	104	303	116	211	62	518	62	734
hBCMA-100	109	262	306	461	212	758	371	1225
hBCMA-30	363	455	281	430	305	606	421	400
FC.	68	160	150	93	391	151	233	237
PBS	311	73	474	152	247	370	870	327

FIG. 35

Evaluation of B cell numbers at treatment day 60 from
 the 12mg/kg (30 ug), 4mg/kg (100ug), and 1.3mg/kg (300 ug) dose of
 hBCMA-Fc groups along with the Fc and PBS control groups.

hBCMA-fc-300					hBCMA-100					hBCMA-FC-30				
Mouse#	%CD4	%CD8	%B220		%CD4	%CD8	%B220			%CD4	%CD8	%B220		
1.0	16.3	11.0	16.4		26.1	14.9	10.1		9.0	2.5	6.9	10.3		
2.0	24.1	11.1	11.6		21.1	11.3	10.6		10.0	13.2	5.2	23.4		
3.0	18.2	7.4	9.9		24.6	13.3	8.3		11.0	15.9	6.4	29.2		
4.0	25.4	13.3	13.1		20.0	11.3	13.4		12.0	14.8	7.6	31.5		
x	21.0	10.7	12.8		23.0	12.7	10.6		x	11.6	6.5	23.6		
sd	4.4	2.4	2.8		2.9	1.7	2.1		sd	6.2	1.0	9.5		
Fc					PBS									
33.0	7.0	8.1	25.4		16.9	8.3	15.5							
34.0	10.7	4.9	15.3		19.1	12.1	19.5							
35.0	18.9	9.3	21.0		7.1	3.4	17.5							
36.0	20.1	11.1	21.0		19.9	11.4	26.5							
x	14.2	8.4	20.7		15.8	8.8	19.8							
sd	6.4	2.6	4.1		5.9	4.0	4.8							

FIG. 36

Specific APRIL binding to Human Cell lines determined by FACS analysis

APRIL binding

HT 29 Colon adenocarcinoma	+	+	+
NCI 460 Lung carcinoma	+	+	+
PC3 Prostate adenocarcinoma	+	+	
C6 Glial carcinoma	+	+	
Raji Burkitt lymphoma	+	+	+
A20 Mouse B cell lymphoma	+	+	+
U266BI Myeloma	+	+	+
A435 Epidermoid carcinoma	--		
A469 Kidney carcinoma	--		
MDA-231 breast adenocarcinoma	--		

FIG. 37
Effect of APRIL, BCMA-Fc and TACI-Fc
truncated on U266BI cell proliferation

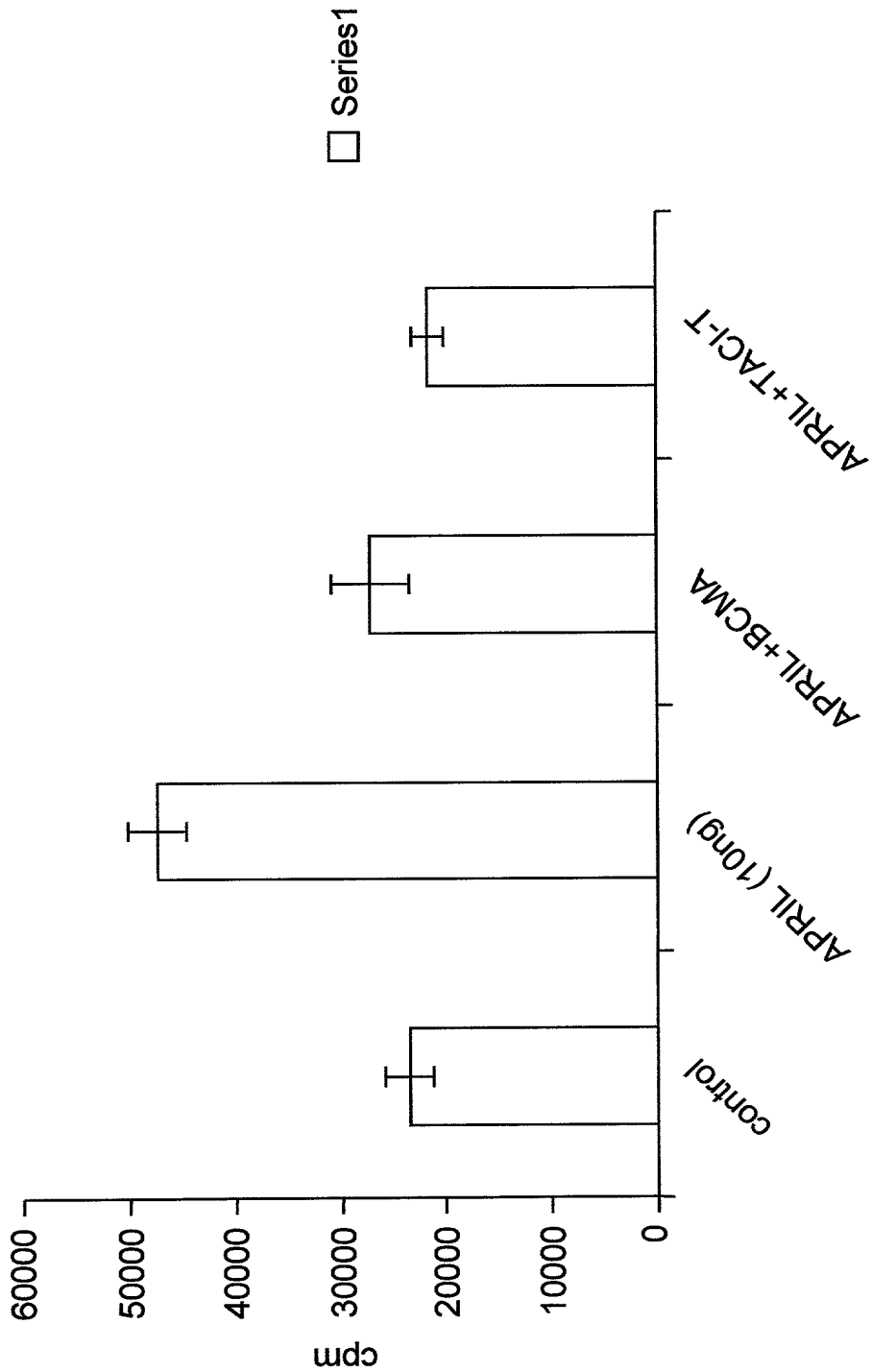


FIG. 38
APRIL and AGP3 stimulates and BCMA-Fc
inhibits B lymphoma cell proliferation

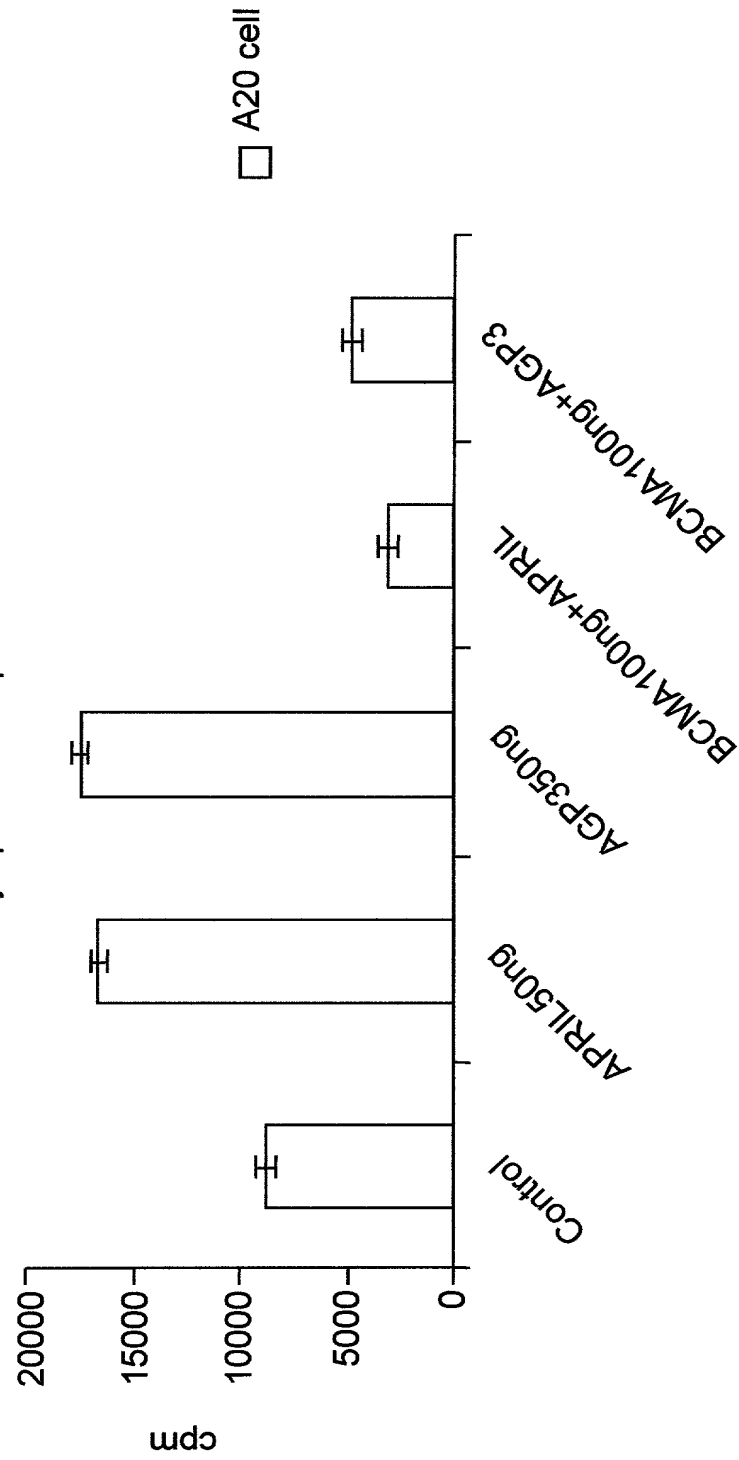


FIG. 39
Effects of BCMA & hTACI on the Growth of A20 in Balb/c Mice

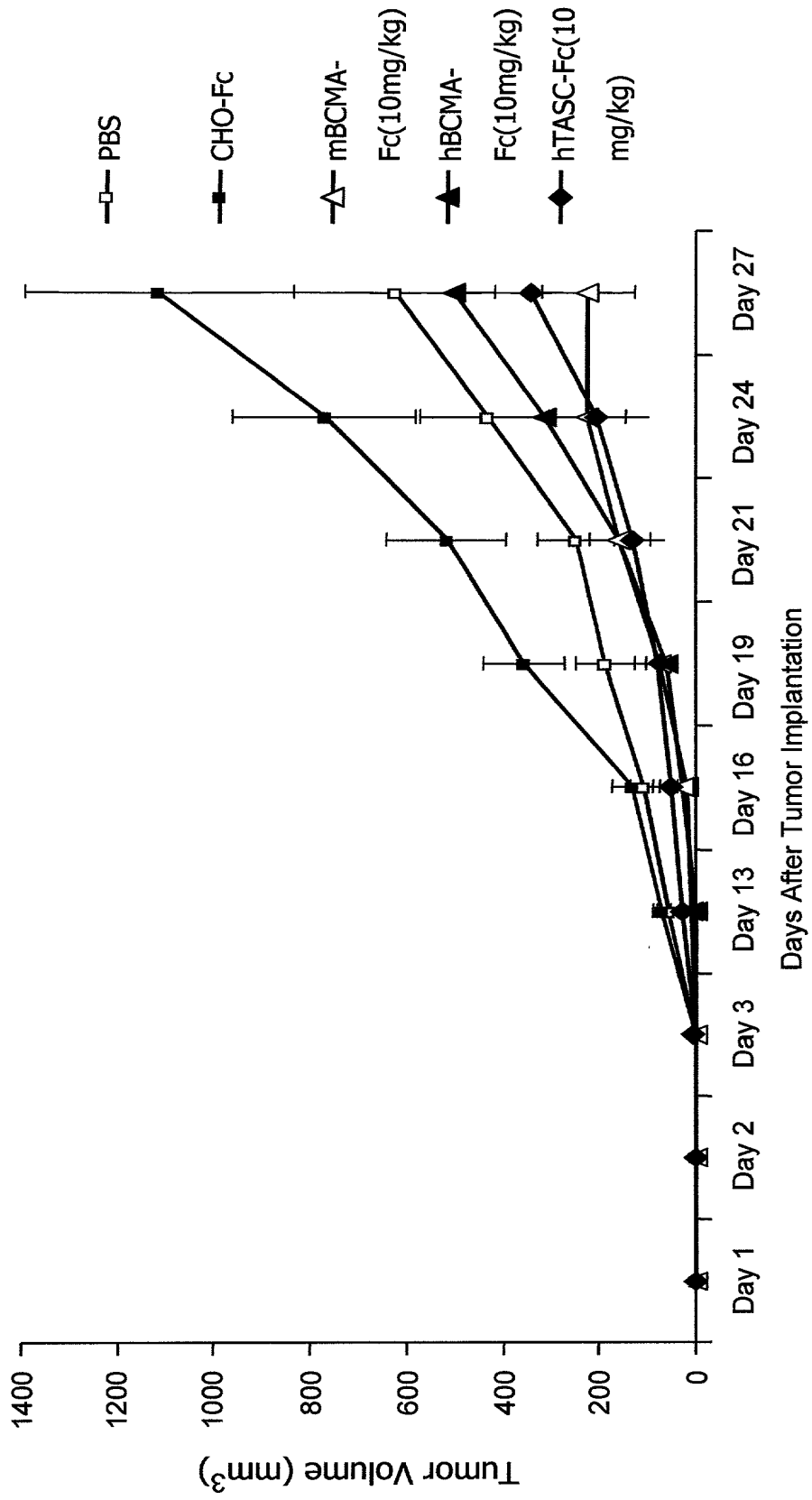
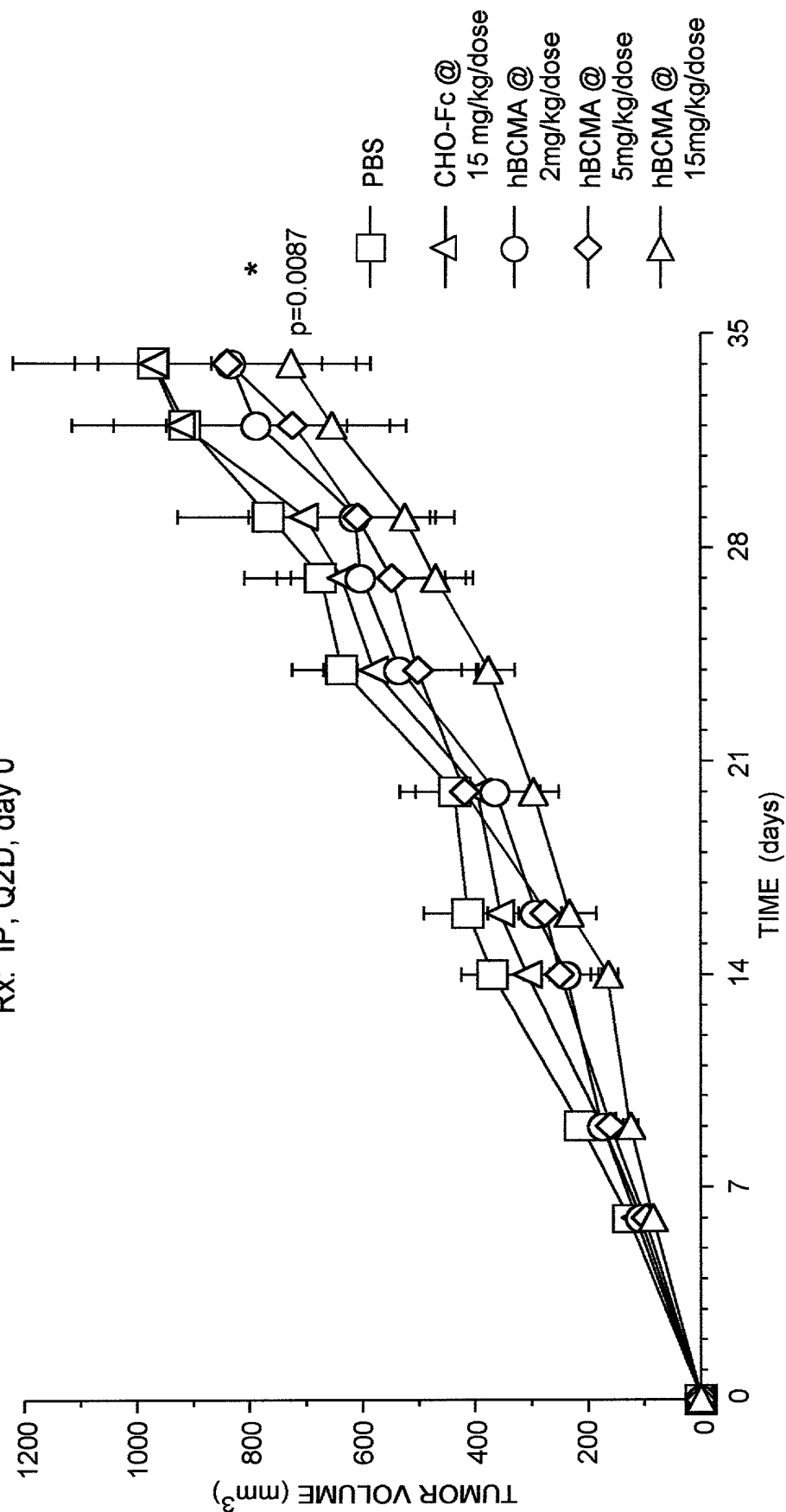


FIG. 40
 EFFECT OF HUMAN BCMA-Fc AGAINST HT-29 SC TUMOR GROWTH
 Rx: IP, Q2D, day 0



* Linear growth ANOVA with Dunnett's correction for multiple testing (n=10/group)

FIG. 41
EFFECT OF MURINE BCMA-Fc AGAINST HT-29 SC TUMOR GROWTH
Rx: IP, Q2D, day 0

